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ZYGADENUS, OR DEATH CAMAS

By

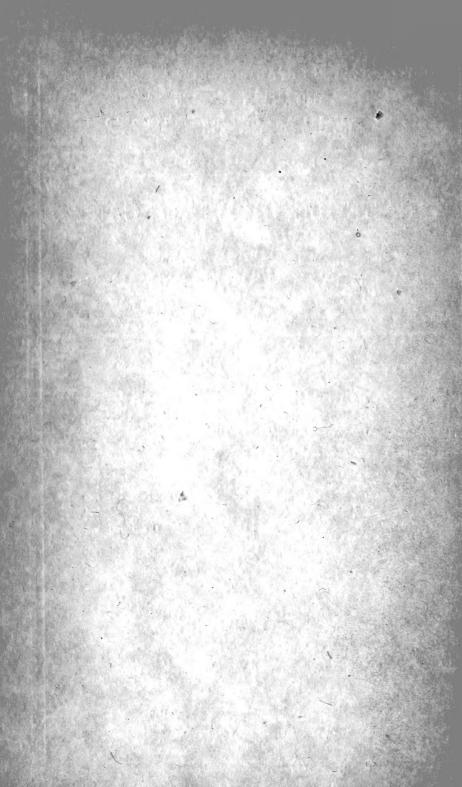
C. DWIGHT MARSH and A. B. CLAWSON, Physiologists and HADLEIGH MARSH, Veterinary Inspector

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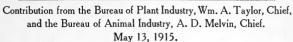
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BULLETIN OF THE USDEPARTMENT OF AGRICULTURE

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(PROFESSIONAL PAPER.)

ZYGADENUS, OR DEATH CAMAS.

By C. Dwight Marsh and A. B. Clawson, Physiologists, Drug-Plant and Poisonous-Plant Investigations, and Hadleigh Marsh, Veterinary Inspector, Bureau of Animal Industry.

INTRODUCTION.

HISTORICAL SUMMARY AND REVIEW OF LITERATURE.

Chesnut and Wilcox (1901, p. 52)¹ say that "the earlier explorers of the Western, and especially of the Northwestern, United States frequently mention the poisonous character of the bulbs of one or the other of the various species of Zygadenus and refer to them as poison camas or poison sego in order to distinguish them from bulbs of two other groups of plants, Quamasia and Calochortus, which were commonly known as camas and wild sego and were much used for food, both by the Indians and by travelers. Accounts of the poisoning of stock from eating the roots and leaves of various species have but recently been sent to this Department."

This statement, perhaps, covers the knowledge of the subject up to that date, although the writers have failed to find much in the way of definite statement among the earlier writers that can be referred to this plant.

In Wyeth's journal of his second expedition to Oregon (Wyeth, 1899) occurs this statement:

16th. Made down the Sandy S. W. by W. 15 miles then 4 S. E. by E. and camped on this stream so far the grass is miserable and the horses are starving and also at last night's camp they eat something that has made many of them sick the same thing happened two year since on the next creek west.

This happened on June 16, 1834, somewhere between Big Sandy Creek and Leckie, in Fremont County, Wyo. The present knowledge of the botany of that region makes it almost certain that the poisonous plant in that place at the time of year mentioned must

¹ For the complete titles of works cited, see the list of literature on pages 45 and 46.

Note.—This paper is intended to supply general information on the relation of Zygadenus to the losses of live stock on the western stock ranges; it is suitable for distribution throughout the western third of the United States.

have been Zygadenus. This is the earliest reference to probable poisoning by Zygadenus which has been found by the writers.

Asa Gray (1848) says of Amianthium nuttallii, now known as Zygadenus nuttallii, "Crescent cum Kamassa esculenta, quo bulbi nocentes viatoribus saepe confusi sunt."

Hooker (1838) says of *Leimanthium nutallii*, which is the same as the species mentioned by Gray, "'Poison or Death Camass' of the Chenooks, from the violent effects of the roots, which create vomiting."

Watson (1880) speaks of Zygadenus venenosus as poisonous and known to the Indians as "Death-Camass," and says, on page 184, that the bulb of Z. paniculatus is also poisonous.

Apparently the Lloyds (1887) were the first to state definitely the symptoms produced by the plant in human beings.

Irish (1889) fed "cammers" to steers without effect.

Hillman (1893) published a newspaper bulletin calling attention to the poisonous character of Zygadenus, and in 1897 he published another newspaper bulletin on the same subject. Also, in another publication (1897b, p. 115), he states that a horse is reported to have been made sick by the seeds of Zygadenus paniculatus in hay.

Coville (1897) says that Zygadenus venenosus causes extreme vomiting and that it is sometimes used by medicine men of the Klamath Indians, mixed with the dried roots of Iris missouriensis and a little tobacco, to give a person a severe nausea, in order to secure a heavy fee for making him well again.

Chesnut (1902, p. 321–322) tells of the knowledge of this plant by the Indians of Mendocino County, Cal., and their use of it for medicinal purposes.

Hunt published an abstract in 1902 announcing the discovery of the alkaloid.

In a copy of McCarthy (1903), apparently annotated by the author, the statement is made that Zygadenus glaberrimus and Z. leimanthoides are poisonous.

Nelson (1906) demonstrated by feeding experiments the poisonous effect of Zygadenus upon sheep.

REVIEW OF PHARMACOLOGICAL WORK.1

The bulbs of Zygadenus paniculatus were found by Collier (1882) to give several alkaloidal reactions, but the first attempts to isolate and determine the chemical and toxic properties of the poison of Zygadenus seem to be those of Reid Hunt,² special expert of the Bureau of Plant Industry in 1901, who worked with the leaves and flowering tops of Z. venenosus. Hunt prepared an alcoholic extract

¹ The review of pharmacological work was prepared by Dr. Reid Hunt, of the Harvard Medical School. ² Hunt's results were submitted in a report to the Department in 1901 and also reported at a meeting of the American Physiological Society. (Hunt, 1902.)

and removed various oily and resinous substances by precipitation with water and extraction with petroleum ether. These resinous bodies were not toxic. Vejux-Tyrode (1904) later obtained similar resinous bodies to which he ascribed a high degree of toxicity, but Heyl and Hepner (1913) could not confirm this.

Hunt purified the extract further, and then, by extraction with chloroform, obtained an amorphous substance alkaline to litmus and giving the usual alkaloidal reactions. It was very slightly soluble in water, but readily soluble in dilute acids. When treated with concentrated sulphuric acid this substance dissolved with the formation of an orange-vellow solution; the color soon became a blood orange, and finally a bright cherry red. This play of colors corresponds almost exactly to that caused by cevadin and to that recently described by Heyl, Hepner, and Loy (1913) for zygadenin, an alkaloid obtained by them from Zygadenus intermedius. Hunt found, as did Heyl, Hepner, and Loy later, that the alkaloid was not readily extracted with ether. When the alkaloid or mixture of alkaloids was further purified, dissolved in alcohol, and the alcohol allowed to evaporate, a clear, glassy residue with a few cubes or prisms was obtained. This began to darken at 185° C. At 197° C. part of it melted to form a red solution, but all of it did not melt until a temperature of 220° C. was reached. It is quite probable that this mixture consisted in part of the alkaloid since isolated by Heyl, Hepner, and Loy and named by them zygadenin. Zygadenin crystallizes from alcohol in "orthorhombic blocks" and melts to a red oil at 200° to 201° C. Hunt pointed out a number of resemblances and also certain differences between the reactions of the alkaloids obtained from Zygadenus and those given by cevadin and other veratrin alkaloids and concluded that both chemically and pharmacologically the two series were closely related.

Torald Sollman, in a report submitted to the Department of Agriculture in 1903, stated that he could find no poisonous principle in Zygadenus other than the alkaloid or mixture of alkaloids found by Hunt.

These results were confirmed by Slade (1905) and by Heyl, Hepner, and Loy. The latter authors carried the work to the point of isolating in pure form an alkaloid which they named zygadenin, although the question whether this may not be identical with some one of the veratrin alkaloids is, perhaps, still open. It may also be doubted whether zygadenin is the most important toxic agent in Zygadenus, for these authors quote Mitchell as reporting that "it (zygadenin) kills guinea pigs slowly and only in comparatively large doses." Hunt found 4 milligrams per kilo of his alkaloidal preparation to be fatal to rabbits in an hour or two. Sollman found about the same amount of cevadin to be fatal.

Hunt found, as did Heyl and Raiford (1911), that the leaves and flowering tops contain more of the alkaloid than the bulbs. performing an extensive series of experiments on animals with the Zygadenus alkaloids, he found that their action was in all essential particulars the same as that of veratrin. They were very irritating to the mucous membranes, as was the powdered plant itself; they produced an intense burning sensation and a very acrid, bitter taste in the mouth; when applied to the skin in alcoholic or chloroform solution, they caused a burning, painful sensation, but the spot later became anæsthetized; they had the typical veratrin effect upon the muscles and, as kymograph experiments showed, affected the respiration, blood pressure, and heart in the same way as does veratrin. It was also shown that the death of animals poisoned with Zygadenus was hastened by attempts to arouse them. This was attributed to the rapid failure of the respiration, circulation, and the muscular system. It was concluded that under laboratory conditions (and probably under field conditions) an essential part of the treatment should consist in allowing the animals to rest.

Hunt also isolated the alkaloids from the urine of animals poisoned with Zygadenus. He found that they were excreted with the urine quite rapidly and demonstrated that under laboratory conditions it was often possible to save the life of poisoned animals (rabbits, sheep, etc.) by the administration of diuretic drugs (caffein, theobromin, sodiosalicylate). Atropin and strychnin seemed to hasten death.

Sollman, after satisfying himself that the toxic action of Zygadenus is identical with that of veratrin, made a study of poisoning by the latter. He found that a single dose often caused prolonged sickness and that small, repeated doses caused no tolerance, but increased the susceptibility, and suggested demulcents, such as linseed decoction, to counteract the corrosive action on the alimentary tract.

It is evident from these chemical and pharmacological studies that the poisonous properties of Zygadenus are essentially those of veratrin, the indications for treatment being the same in the two cases.

DESCRIPTION OF ZYGADENUS.

The genus name Zygadenus is used in this paper as defined in Robinson and Fernald's revision of Gray's Manual and in Coulter and Nelson's New Manual of Botany of the Central Rocky Mountains. It includes the three genera, Zygadenus, Anticlea, and Toxicoscordion, of Britton and Brown's Illustrated Flora. The plants are erect, perennial, glabrous herbs, growing from a rootstock, or, as in the case of all the western species, from a tunicated bulb, with a leafy stem. The leaves are grasslike, long, narrow, and keeled. The flowers are greenish yellow or white, borne in a terminal raceme or panicle. This

raceme varies in the different species from an almost solid head, as seen in Plate I, to a very loose, elongated panicle, there being a considerable range of variation in the inflorescence within the limits of the same species. The perianth is spreading, withering-persistent, the sepals bearing one or two glands near the base. The stamens are free or attached to the bases of the segments. The capsule is three lobed and dehiscent to the base in maturity.

The species of Zygadenus are spring and summer plants. On May 8, 1913, in the neighborhood of the Greycliff station, Mont., Z. venenosus was about 4 inches high, the largest plants not exceeding 6 inches, and the flower scape was not visible. On May 11 the plants were in bud, and they blossomed through the month of June. Seeds were formed the last of June and early in July, and after the middle of August the plants had largely disappeared.

Zygadenus elegans was in full blossom near Red Lodge, Mont., on July 20, at an altitude of approximately 6,000 feet. In 1910 Zyga-denus coloradensis was in blossom in Colorado at about the same

time (July 20) at an altitude of about 10,000 feet.

In Montana, Zygadenus venenosus grows typically at lower levels than Z. elegans. As stated by Chesnut, its favorite habitat is in the shallow ravines occurring on hillsides. It does not grow abundantly on dry hillsides nor in wet ravines, but it is very commonly found in the shallow depressions on the north slopes of bench lands. Z. elegans grows at higher levels (Rydberg gives as its limits 6,500 to 12,500 feet) and in locations where more water is available than is necessary for Z. venenosus. While it grows readily on hillsides, it reaches its best development in size in distinctly wet places, sometimes immediately in contact with rivulets.

In California and Oregon, Zygadenus venenosus grows in the meadows, while Z. paniculatus grows upon the hillsides. Z. venenosus is more common on north slopes and Z. paniculatus on south

slopes.

The species of Zygadenus may grow as more or less scattered individuals, but sometimes they are massed together in large areas, including, perhaps, several acres, in which, at the time of flowering, they seem to be the principal vegetation and give a characteristic greenish yellow color to the landscape.

The species of Zygadenus are distributed very widely in the United States and are found as far north as Alaska. They occur most abundantly from the Rocky Mountains west to the Pacific, and their importance as stock-poisoning plants is confined almost entirely to this part of the United States. Plate I shows the general appearance of Zygadenus venenosus. This is an Oregon plant and shows the flowers as they appear at the beginning of blossoming. Later, the raceme is

more extended, as shown in Plate II, which is reproduced from a photograph of a Montana plant. Plate III shows the plant after the seed is formed.

COMMON NAMES OF ZYGADENUS.

The species of Zygadenus are known under a large number of popular names. The most common perhaps is death camas. In the Northwest perhaps lobelia is the name used even more generally than death camas. Other names are soap plant, alkali grass, water lily, squirrel food, wild onion, poison sego, poison sego lily, mystery grass, and hog's-potato. Z. glaberrimus is said to be called cow-grass.

POISONOUS SPECIES OF ZYGADENUS.

The following species of Zygadenus are said to be poisonous: Z. elegans, Z. falcatus, Z. fremontii, Z. glaberrimus, Z. intermedius, Z. mexicanus, Z. nuttallii, Z. paniculatus, Z. venenosus.

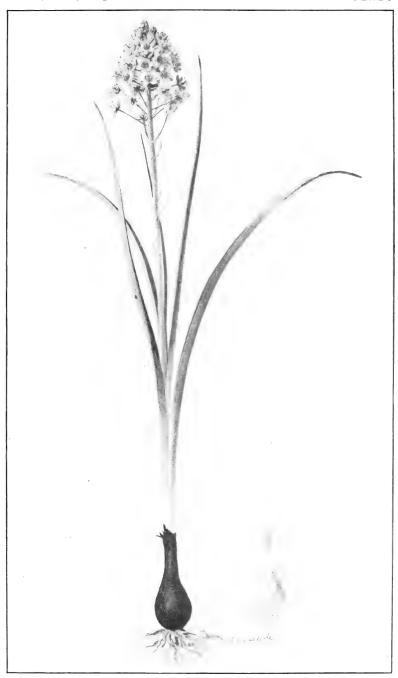
This list is given in accordance with the statements of various authors, and no attempt has been made to revise it from the standpoint of the systematic botanist. Apparently all species of this genus may be presumed to be poisonous. Even Zygadenus coloradensis, which has been shown not to be injurious to stock in Colorado, has the same poisonous principle as the other species, but in smaller quantity.

LOSSES OF LIVE STOCK BY ZYGADENUS.

As already stated, there is reason to think that deaths of cattle and horses from Zygadenus poisoning are not numerous. With sheep, however, the losses are very heavy, but it is impossible to make even an approximate estimate of these losses. It is probable that they are much greater than is generally supposed, for in the sheep-grazing regions many, perhaps most, of the herders do not know the plant and consequently do not recognize it as the cause of illness and death in the bands under their charge. The lupines, without any doubt, have been blamed for many of the cases of poisoning by Zygadenus.

Chesnut and Wilcox (1901, p. 53) state that 636 sheep died from Zygadenus poisoning in Montana in 1900 and that 3,030 were poisoned. In one locality in Wyoming 500 sheep died out of a total of 1,700 poisoned, and in one county it was said that 20,000 died in 1909. The writers of this paper investigated a case in Montana in which 500 sheep died within a few hours, the probable cause being Zygadenus.

There is no doubt that this plant is one of the sources of heaviest loss to sheep owners, especially in Wyoming and Montana. There is good reason, too, for thinking that many of the losses in Oregon, Utah, and California which have been ascribed to other plants were really caused by Zygadenus.



Zygadenus Venenosus from Klamath Agency, Oreg.



ZYGADENUS VENENOSUS FROM MONTANA, IN BLOOM.



ZYGADENUS VENENOSUS FROM MONTANA, IN FRUIT.

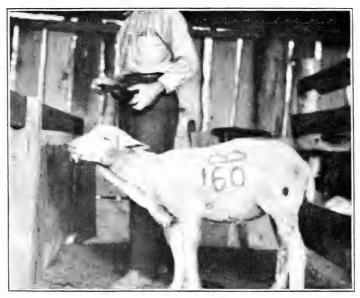


Fig. 1.—Sheep No. 160, Showing Salivation and Position Indicating Nausea.

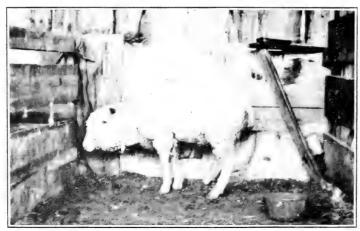


Fig. 2.—Sheep No. 192, Showing Salivation and Attitude Indicating Nausea and General Discomfort.

ANIMALS POISONED BY ZYGADENUS.

Swine are said to eat Zygadenus bulbs with no bad results (Parsons, 1904, p. 8).

Cattle are susceptible to the poison and there are reports of resulting deaths. So far as has been learned, however, deaths of cattle from this cause are not common, and it is not especially to be feared

by the cattlemen.

Many cases of horses poisoned by this plant are reported. The animals are made very sick, but apparently most of them recover. Mr. Uttermohl, of Bigtimber, Mont., who has had considerable experience with Zygadenus, is of the opinion that some of those that

recover are permanently injured.

Sheep are the animals most frequently poisoned. This is probably due in part to a greater susceptibility to the toxic principle of Zygadenus, but very probably it is to a considerable extent due to the way in which sheep are managed upon the range. While grazing, they are frequently herded rather compactly, so that they eat the forage closely, and when passing over a Zygadenus area many of them may eat a large quantity of this plant.

The cases of human poisoning are mostly of children, who find the bulbs attractive and sometimes collect them instead of the edible camas, species of Calochortus and Camassia. Most of these cases

recover, but there have been a number of fatalities.

SYMPTOMS PRODUCED BY ZYGADENUS POISONING.

With the exception of the work of Chesnut and Wilcox, nothing has been published in regard to the symptoms exhibited by grazing animals. These authors (1901, p. 61) state that the principal symptoms of poisoning in sheep are salivation, nausea, uneasiness, staggering, muscular incoordination, paralysis, and convulsions. The animals sometimes lie many hours before death. The writers mentioned state also that cattle and horses have spasms.

Several investigators have mentioned some of the symptoms in man. Heller (1909, p. 52) gives the symptoms (quoting from Dr. Lee, of Carson) as "nausea, headache, followed by more or less stupor." He states that the heart's action was lessened in frequency, while the strength of the pulse remained normal. The respirations were almost normal. In another case vomiting was followed by the loss of all power of feeling.

Heyl and Raiford (1911, p. 64) and Hunt (manuscript) speak of the irritating character of the dust when the dry plant is being ground, which leads to sneezing on the part of those doing the work.

The Lloyds (1887) give as symptoms in man "extreme thirst, constant vomiting, dilation of the pupil, coma, and inflammation of the stomach." They also say that one case had very violent convulsions.

Chesnut (1902, p. 321) says that the symptoms in poisoned Indians are "burning and smarting in the mouth and esophagus, dumbness, nausea, profuse vomiting, foaming at the mouth, dizziness, and mania."

Mitchell and Smith (1911) experimented with the extract on guinea pigs, both by subcutaneous injection and by feeding per os, and found salivation, vomiting, excitement, paralysis (first of the hind legs), rapid respiration becoming slow and labored, heightened reflexes, spasms, heartbeat slowed, and death, under fatal dosage, in 20 to 30 minutes. When injected into dogs under anæsthesia, the general effect was to reduce the rate of heartbeat and respiration and to produce marked intestinal peristalsis. The heart stopped before the cessation of respiration.

Hunt, Vejux-Tyrode, and Mitchell and Smith experimented on frogs, producing paralysis, which showed itself in an inability to draw up the legs readily after extension. Hunt considers that it produces an effect directly on the muscles as well as on the central nervous system.

Chesnut and Wilcox (1901) and Hunt (manuscript) experimented with rabbits, Hunt stating that the rabbits exhibited salivation, nausea, muscle changes, heightened reflexes, and convulsions.

Summarizing the published statements in regard to the symptoms of Zygadenus poisoning, it may be said that the most evident symptoms in the higher animals are salivation, nausea, more or less complete paralysis, reduced rate of heartbeat and respiration, and convulsions. The results on frogs are not so marked, as would be expected from the less complicated nervous system, and the principal thing noticed apparently is paralysis.

GENERAL STATEMENT OF EXPERIMENTAL WORK.

Experimental work upon Zygadenus has been carried on for five seasons, in 1909 and 1910 at Mount Carbon, Colo., and in 1912, 1913, and 1914 at Greycliff, Mont. Table I gives a summary of these experiments. In 1909 six head of cattle were fed experimentally on Zygadenus coloradensis (Table I, section A). In 1910 a steer and four sheep were fed (Table I, sections B and E). In 1912 there were 18 cases of experimental feeding of Zygadenus venenosus to sheep (Table I, section F). In 1913 Zygadenus venenosus from the neighborhood of the station was fed to 61 sheep. In this section of the table are also given the results of one experiment in feeding Zygadenus venenosus from the Stanislaus National Forest, Cal., to a sheep. In 1913 Zygadenus elegans, collected near Red Lodge, Mont., was fed to 6 sheep (Table I, section H). In 1914 there were 110 cases of feeding of Zygadenus venenosus to sheep (Table I, section I) and five experiments of feeding to sheep Zygadenus elegans from the

Fishlake National Forest, Utah (Table I, section J). There were also two experimental feedings of *Zygadenus paniculatus* from Ephraim, Utah (Table I, section K). Three head of cattle in 1913 were fed upon *Zygadenus venenosus* (Table I, section C), and a horse was fed twice upon *Zygadenus venenosus* and once upon *Zygadenus elegans* (Table I, section D).

So far as possible, the feeding experiments were carried on under natural conditions. To this end the animals were, ordinarily, deprived of food for about 24 hours, and then the plant to be tested was offered to them. If they did not eat readily, they were tempted by mixing the plant, sometimes ground up, with hay or grain. As it was difficult to get any large number of cases by feeding, on account of the dislike of the animals to the plant, resort was had to drenching and forced feeding. In the drenching experiments, the plant was ground and suspended in sufficient water to make the administration possible, the drenching being done in most cases with the animal upon its haunches. Forced feeding was conducted in some cases by placing the plant by hand, a little at a time, in the animal's mouth. In the majority of experiments in forced feeding, however, a veterinarian's ordinary balling gun was used, and the ground material was fed as fast as the animal would swallow it.

The terms under "Severity of illness" are used in the following way:

"Not sick" includes cases in which no symptons appeared.

"Symptoms" includes cases in which there was slight salivation for a few minutes, some regurgitation, some licking of the lips, indicating nausea, or indications of uneasiness.

"Slightly sick" includes those in which salivation was continued for an hour or more, with considerable regurgitation. Depression, slightly labored respiration, and temperature reduction may occur.

"Sick" includes cases exhibiting vomiting, weakness, and sometimes hypersensitiveness and trembling.

"Very sick" cases were characterized by prostration, extreme respiratory difficulty, and subnormal temperature.

47698°-Bull, 125-15-2

Table I.—Summary of feeding experiments with Zygadenis in 1909 and 1910 at Mount Carbon, Colo., and in 1912, 1913, and 1914 at Greycliff,

[Animals designated by an asterisk (*) preceding the serial number were given dry material; but the weight is stated in the form of an equivalent amount of green plant, the loss in drying having been determined by a number of experiments.]

BULL		125, U. S.			0.				Œ.		
Location from	was obtained.			North of camp.	Station. Do.	Do.	В о. Во.	Beartooth National Forest.	North of camp.	Do. ·	Do.
Pounds used per	weight of animal.	Per 1,000 pounds. 8.31 3.16 13.16 7.92	18,14	325	58	111	12.1 12.9	15	Per 100 pounds. 16.7	82	63 40.7
1,550	Mesun.				Recovery.	do	* * * * * * * * * * * * * * * * * * *				Recovery.
c	remeay.	None do	do	ор	op op	ор••••	ор	op	ор	do	op op
	Severity of himess.	Not stek 40 40 40 40 40	do	ф	Symptoms	Symptoms after eating 46% pounds.	Not sick	do	op	do	Leaves, stems, and fruit. Leaves, stems, and fruit. Symptoms Go Recovery
Part of plant used (fed	unless otherwise stated).	Leaves and buds Bulbs (in chop) Buses and buds Buths (in chop) Geaves, stems, and	Leaves, stems, flowers, and bulbs.	Leaves, stems, fruit, and bulbs.	Leaves. stems, and	nowers.	Leaves of Z.venenosus. Leaves, stems, and flowers of Z. vene-	nosus. Leaves, stems, flowers, some fruit, and bulbs of Z. elegans.	Leaves, stems, flowers,	Leaves, stems, flowers,	Leaves, stems, and fruit.
	Date of experiment.	July 14 to 16. July 15. July 17 to 19. July 17 to 19. July 19 and 20. July 20.	Λug. 1 and 2	1910. Aug. 8 to 13	1913. June 6 and 7. June 12 to 17.	June 21 to 27	June 15 to 18	July 24 to 29	1910. July 15 to 19.	July 24 to Aug. 5	Aug. 15 to 24Aug. 25 to 31
Woight of	plant used.	Pounds. 5. 07 1. 87 6. 58 5. 94 12. 22	7.62	97.5	29	55.5	9.5	10.5	, 16	83	45. 5 39. 5
	Weight.	Pounds. 610 620 500 750 610	420	300	±500 ±500	7200	·	₩ 200	100	100	72 97
Animal.	Designation.	A.—Cattle given Z. coloradensis: No. 112 No. 117 No. 113 No. 113 No. 113 No. 113 No. 115	No. 107	B.—Steer given Z. coloradensis: No. 119	C.—Cattle given Z. venenosus: No. 655. No. 660.	No. 659.	D.—Horse:	No. 125	E.—Sheep given Z. coloradensis:	No. 112	No. 111 No. 118

George Hughes's.	Do.	Do.	Do.	Do.	Do.	Do.	Do.	Do.	Do.	Do.	Do. Do.	Do.	Do.
1.134	8, 461	18.831	10,377	1, 762	2.144	1.632	1.182	. 853	. 929	. 560	. 141	. 264	.172
			***************************************	Recovery .	do	Recovery.	do	Death	Recovery	Death	Recovery Death	Recovery	Recovery .
ф	ор	ор	do	do	до	Potassium permanga- nate; whis-	Caffein sodio- benzoate.	do	None	Ammonia; caffein so-	diobenzoate; tannic acid. Nonedo Potassium permanga- nate; cafein	ate; whisky; digitalis. None Caffein sodio- benzoate; potassium	nate.
Not sick	do	ор	ор	Sick	Slightly sick after tak- ing 0.79 pound; very	Sick. Not sick. Slightly sick after tak- ing 0.33 pound; very sick.	Slightly sick after tak- ing 0.771 pound;	Sick after taking 0.33	Symptoms after tak- ing 0.33 pound,	Sugnery Sick. Death	Symptoms Sick Death.	Sick from Zygadenus; killed in drenching with potassium per- manganate	Symptoms
Leaves, stems, and	Leaves, stems, flow-	Leaves, stems, flow- ers, and fruit; partly	Leaves, stems, flow-	Leaves, stems, and	run (arenen).	Water control (drench) Leaves, stems, and fruit (drench).	do	Seed heads (drench)	Leaves and stems (drench).	Seed heads (drench)	dododo	dodo.	op
1912. June 25 to 27	June 26 to July 4	June 26 to July 7	do	July 6	July 7	July 10.	July 11	July 13	do	July 14	July 17. July 18. July 19.	July 20. July 23.	July 24
.771	5.5	14.5	5.5	1,322	1,565	1,102	1,212	. 661	. 771	.308	. 110 . 121 . 275	.198	.143
89	65 to 57½	77 to 71	53 to 45	75	73	77 67 <u>3</u>	102½ to 84½	773	83 to 75½	55	78 51 <u>3</u> 92	75	88
Z. venenosus:	No. 163	No. 175	No. 181	No. 176	No. 159	No. 172 No. 168	No. 162	No. 161	No. 171	No. 174	No. 172 No. 160.	No. 176 No. 181	No. 175.

Table I.—Summary of feeding experiments with Zygadenus in 1909 and 1910 at Mount Carbon, Colo., and in 1912, 1913, and 1914 at Greycliff, Mont.—Continued.

Location from	was obtained.	George Hughes's	Station.	Do. Do.	Do.	Do.	Do.	Do.	Do.	Do.	.00	Do.	Do.	Do.	Do.		Do. Do.	Do.	Do. Do.	Do.
Pounds used per stated	weight of animal.	Per 100 pounds.	986	.595	.612	1.905	.746	3.519	.586	000.	200	.585	.555	.568	.572		1.397	.546	1.536	.384
Resnit		Recovery		Recovery.	do	do	Death		Recovery .	9	000	Recovery.	do	do	Death		Recovery . Death, due	phin.(?) Death	Recovery .	Death
Remedy		Nome	do		None	do	Strychnin; gin.	None	do do Dotossium nor	manganate.	fein citrate.	None Diuretin; caf-	None	Diuretin; caf-	tem curate. Diuretin; caf- fein citrate:	gin; am-	None Eserin; strych-	phin. Apomorphin; e s e r i n ;	strychnin. None Eserin; strych-	Gin Death
Severity of illness		70,00	Not sick do	Sick Very sick	Slightly sickdo	Sick	Death	Not sick	Sick	do	000	Not sick Very sick	do	Sick	Death		Sick Death	do	Very sick Sick	Death
Part of plant used (fed	stated).	Toorses	do	Bulbs (drench)	Bulbs (drench)	Leaves	Leaves (drench)	LeavesBulbs (in alfalfa)	Leaves (drench)	do		Leaves (in alfalfa) Leaves (drench)	Leaves, stems, and	Duds (arench).	ор		Leaves	Leaves (drench)	1 .	and Howers (drench). Leaves (drench) Death
Date of exnariment	Date of experiments.	1913. More 11 cmd 19	May 11 to 13. May 12 and 13	May 13. May 14.	May 15	May 15 and 16.	May 17.	May 17 to 20	May 19.	Моч 99	May 26	May 22 to 24	May 24	do	May 26		do May 26 to 31	May 27.	May 28 to 31	238 May 30.
Weight of	plant used.	Pounds.	.631	441	.441	1.219	.485	220	.375	976	017.	.441	.386	.551	.392		.403	.430	.883	. 238
	Weight.	Pounds.	64 64 66	74 59 to 61	72 64 to 64	64 to 613	74 00 71	71 to 623	64 to 583	04 to 004	\$04 01 504	71 to 63½ 77½ to 72½	69½ to 67½	97 to 91½	683		70½ 71	त्त्र ⁴ 00 त्र	573 to 53 71 to 673	62
Animal,	Designation.	G.—Sheep given Z. venenosus:	No. 191	No. 193 No. 187	No. 195	No. 198	No. 189	No. 190	No. 184	No 305	INO. 200	No. 200.	No. 206	No. 203	No. 197		No. 198.	No. 188	No. 191	No. 192

Do.	Do.	D0.	Do. 00.	Do.	DO	0 D0.	Avery, Cal. Station.	D0.	Do.	Do.	Do. Do.	Do.	Do.	Do.	Do.	0.000	
.385	.338	.430	.386 .385 .435	.496	. 495 . 550 . 550 3. 695	.325 1.728 1.789	2.139	1.672	.633	5.896	2.800 3.956	2.371	2.896	3,155	5,597	. 092 . 151 . 110	
Recovery .	Recovery.			Recovery.	Death	Recovery.	do	0 0					Recovery.	Recovery.	qo	dodo	
Eserin; strych- Recovery	Nonedo	Tannic acid 1	None Tannic acid 1 None.	Tannic acid 1	None Tannic acid 1 None Tannic acid 1	None do Tannic acid;1 Ensom salts	ZE	4	op	ор	do do	ф	Diuretin	None Tannic acid; diuretin.	None	00 00 00 00	
Sick	Not sick Symptoms	Not sick	op op	Slightly sick	Sick. Death Sick	Not sick Sick do	Symptoms.	Not sick	do Symptoms.	Not sick	op op	op	Sick	Not sick Sick	ор.	Symptoms Not siek. Symptoms after eating 45 grams.	
do	Leaves. Leaves, stems, and	Leaves, stems, and	Η	Leaves, stems, and	do do Leares, and Houses, stems, and	dodo.	Leaves (in bran) Leaves, stems, and flowers.	do.	do. Leaves, stems, flowers, and young fruit (in	bran). Leaves, stems, and	do Leaves, stems, and	Young seed heads (in	Leaves, stems, and	do Seed heads with a few leaves and flowers	Seed heads in bran	Seed (in bran)do.	1 Drench.
do	June 3 to 5	June 4	June 5 June 5 to 7	June 6	June 7 June 7 June 7 to 10.	June 7 to 11	June 13 to 19	June 15 to 18.	June 20 to 22. June 22 to 28.	June 23 to 30	June 26 to 30	July 4 to 7	July 6 to 8	July 6 to 11. July 6 to 12.	July 16 to 19	Aug. 13. Aug. 13 and 14. Aug. 14. Aug. 15 and 16.	
.322	1.146	.298	.293	.399	.392 .516 .505 3.307	. 220 1.455 1.146	3.274	1.213	5.247	4.894	2. 282 3. 086	1,731	1.896	3,472	4.112	.055 .110 .066 .154	
833	65 to 581 631	\$69 to \$69	79 to 73‡ 50½ to 52	80½ to 77½	794 to 754 932 to 90 913 895 to 78	673 to 593 843 to 71 64 to 57	85½ to 77 71 to 66	72½ to 65 78 to 63	73 to 69	83 to 67	81½ to 67 78 to 62½	73 to 63½	89 to 76½	84 to 70 110 to 98½	73½ to 66	73 to 70 64 to 58	
No. 195	No. 196	No. 206	No. 200 No. 201 No. 199	No. 210	No. 209 No. 212 No. 213 No. 208	No. 207 No. 211 No. 214	No. 216. No. 193	No. 202 No. 215	No. 201	No.210	No. 209 No. 196	No. 190	No. 216	No. 195 No. 203	No. 193	No. 199 No. 196 No. 218 No. 185	

Drench.

Table I.—Summary of feeding experiments with Zygadenus in 1909 and 1910 at Mount Carbon, Colo., and in 1912, 1913, and 1914 at Grapcliff, Mont.—Conti nued.

Date of experiment.
Aug. 16 and 17. Seed (in bran)
July 25 and 26 Leaves, stems, flowers,
July 25 to 31 Leaves, stens, flowers,
Leaves, stems, flowers,
July 28. Leaves, steins, flowers, and some fruit
July 29
May 27 to 29 Leaves, stems, flowers, and buds (With
May 27 to 30 Fruss). May 28 Leaves, stems, flowers, and buds (forced feeding).
May 28 to 30 Leaves, stems, flowers,
and buds. Leaves, stems, flowers, and buds (with
May 29. Leaves, stems, flowers, and buds (forced
.421do .

Do.	Do.	D0.	Do.	00. 00.	Do.	0. 0. 0.	Do.	Do. Do.	Do.	Do.	Do.	0000 0000 0000	Do. Cabin Corral.	
. 469	. 499	.550 .551 1.643	. 518	. 518 . 529 . 529 1. 236	. 529	. 529 . 529 . 622	.590	. 420	. 662	. 359	.757	. 770 . 826 . 826	. 805	
	Recovery.	Recovery.	Recovery.	op op		Recovery.	do		Recovery.	do	Recovery.	Recovery . do	do	
ор	ор	op	Tannic acid 1 Recovery	None Tannic acid 1 None do	do	do	None	None	Tannic acid	None	ор	dododododododod	None dodo	² Very few grams.
do.	Symptoms	do. Not sick (?)	Sick	do Slightly sick Sick. Not sick.	do	do do Sick	Very sick	Not sickdo	Slightly sick	do. Not sick	Symptoms	Not sick. Symptoms. do. Slightly sick.	Not sickdo	2 Very f
Leaves and some young buds (with	Leaves and some S young buds (forced	do d	flowers (with grass). Leaves, stems, flowers, and buds (forced		grass). Leaves, stems, flowers, and buds (forced	leeding). do. do.	Leaves, stems, flowers, and buds (with	grass). do Leaves (forced feed-	ng). do	do Stems, flowers, and young fruit (forced	feeding). Leaves, stems, flowers, and young fruit	(forced feeding). do. do. do. do.	do. Leaves, stems, and flowers (with grass).	
441 do	ф	June 2do. June 2 and 3	June 3	.do. June 4 June 4 and 5.	June 5	June 6 June 8	June 9	June 15	do.	June 16.	June 17	do. June 18. do. June 19.	June 20.	¹ Drench.
. 441	. 529	.465 .457 1.323	. 539	. 526 . 529 . 503 1. 323	. 476	. 471 . 479 . 666	.649	. 441	.612	. 269	. 628	. 639 . 851 . 843 . 789	. 784	
94 to 85½	106 to 98	84½ to 78½ 83 to 78 80½ to 76	104 to 100½	101½ to 89½ 100 to 97½ 95 to 87¾ 107 to 101	90 to 84	89 to 81 90½ to 81½ 107	110 to 1023 98 to 93	105 to 109 91 to 84	92½ to 87	85 to 79 75 to 71	83 to 79	83 to 74 103 to 95 102 to 95 79½ to 75	79 to 75 109½ to 99½	
No. 248	No. 267	No. 239 No. 256 No. 243	No. 229	No. 235 No. 245 No. 257 No. 231	No. 265	No. 249 No. 266 No. 264	No. 268	No. 247	No. 269.	No. 282	No. 237	No. 238 No. 235 No. 245 No. 239	No. 256	

rench.

Table I.—Summary of feeding experiments with Zygadenus in 1909 and 1910 at Mount Carbon, Colo., and in 1912, 1913, and 1914 at Greycliff, Mont.—Continued.

Management of the Control of the Con	Location from	which plant used was obtained.	Cabin Corral,	D0. D0.	Do. Do.	Do. Do.	Do.	Do.	Do.	Do.	.Do.	Do.	Do.	Do.
	Pounds used per	stated weight ofanimal.	Per 100 pounds. 1.213	. 992 . 992 1.188 1.436	1.434	1.871	1,983	1,984	2, 205	1.103	2, 205	2.756 2.756	2, 156 1, 103	.662
		Result.	Recovery.	ор. ор.	Recovery.	ob		Recovery.	do		Recovery.	do		
		Remedy.	Tannic acid (in capsule).	Nonedo	Η		(In capsule).	qo	Tannic acid (in capsule).	None	ор	Tannic a c i d	Nonedodo	dodo
		Severity of illness.	Slightly sick	Sick do	Not sick	do. Slightly sick	Not sick	Sick	Slightly sick	Sick	Slightly sick	dodo.	Not sickdo.	do
	Part of plant used (fed	unless otherwise stated).	Leaves, stems, and flowers (forced feed-	ing). do. do. Leaves, stems, flowers, and buds (forced	reeding).	op	Leaves, stems, flowers, and buds (in 7 forced	feedings). Leaves, stems, flowers, and buds (forced	Leaves, stems, and flowers (forced feed-	Leaves, stems, and	Leaves, stems, and	dodo	Leaves, stems, and	dodo
		Date of experiment.	June 20	.dodododoJune 22	do. June 23.	June 24	ор	do	June 25	do	ф	June 26do.	June 27do	op
	Woicht of	plant used.	$Pounds.\\1.043$. 838 . 843 1. 093 1. 300	1.362	1.413	2, 033	2, 202	2, 282	1.142	2, 183	2, 205	$\frac{2,425}{1,064}$. 655
		Weight.	Pounds. 86 to 81	84½ to 79 85 to 78 92 to 88½ 90½ to 86½	95 78 to 70½	75½ to 70½ 112½ to 98½	$102\frac{1}{2}$	111 to 101½	103½ to 98½	103½ to 99½	99 to 94½	80 to 78 82 to 81	112½ to 100 96½ to 95½	66
	Animal.	Designation.	L—Sheep given Z. venenosus—Con. No. 234.	No. 249 No. 265 No. 266 No. 251	No. 263 No. 232	No. 253 No. 229	No. 264	No. 268	No. 240	No. 250	No. 280	No. 262 No. 278	No. 247 No. 255	No. 277

.441 Station.	00. 00.	Do. Do. Cabin Corral.	Do. Do.	Station,	Cabin Corral,	Do.	Do. George Hughes's.	Do.	Do. Do.	Do. Do.	Do.	Do.
. 441	. 881 1.103 1.102	1, 322 1, 323 1, 442	1.432	1, 543	1,543	1.763	1.764	. 994	. 992	. 992	.331	1.653
	Recoverydo Killed by choking on cap-	Sute. Recovery do	Recovery.	Recovery.	do	do	do	Recovery.	do	do Death Recovery.		Recovery.
ор	do do Tannic a c i d (in capsule).	do None Tannic a c i d (in capsule).	Nonedo	ор	Tannic a c i d (in capsule).	Tannic a c i d (in capsule);	None Tannic a c i d (in capsule); oil and tur-	Tannic a c i d	None Tannic acid; 1 caffein sodio-	Nonedo Tannic acid	None	Potassium per-manganate.
do		do Not sick.	Slightly sick Not sick	Symptoms	do	Very sick	Sick. Death	Sick	do.	Death. Sick.	Symptoms	dod
Seed heads, nearly developed (forced feed-	Ing., Slightly sick do	do do Seed heads, some fully develope d and others half devel- oped (forced feed-	do. Leaves, stems, flowers,	Seed heads, v e r y young (forced feed-	Seed heads, half developed (forced feed-	dodo.	Seed heads, fully developed (forced feeding).	do	op	do do. do.	Seed heads, fully de-	veloped (drefich), Seed heads, fully de- veloped, mixed with potassium perman- ganate (drench).
.377 June 29	. do. June 30. do.	July 1do July 2.	July 2 and 3	July 3	do	July 5	July 7.	do	July 8		July 10	do.
.377	. 679 1. 053 1. 047	1.177 1.164 1.024	1.182	1,582	1.628	1.640	1.508	1.047	$\frac{1.002}{1.111}$	1.295	.326	1.554
851	77 to 64 95½ to 90½ 95	89 to 833 88 to 823 71 to 683	82½ to 76½ 115½ to 98½	102½ to 101	105½ to 98½	93	85½ to 80 83½	105½ to 100½	101 to 95 112 to 111½	130½ to 126½ 87 86 to 79	98½ to 92	94
No. 243	No. 257	No. 237 No. 238 No. 281	No. 283	No. 235	No. 267	No. 282	No. 239	No. 245	No. 269	No. 233 *No. 249 *No. 251	No. 246	*No. 266

Drench.

Table I.—Summary of feeding experiments with Zygadenus in 1909 and 1910 at Mount Carbon, Colo., and in 1912, 1913, and 1914 at Greycliff, Mont.—Continued.

Location from	which plant used was obtained.	George Hughes's.	Do.	Do.	Do. Do.	Do. Do.	Do.	Do.	Do. Do.	D0. D0.	Do.	Do.
m =	statêd weight ofanimal.	Per 100 pounds. 0.441	. 441	.440	. 440 . 759 . 881	. 882	.881	.740	.881	. 882 . 665 . 882	.882	. 882
	Result.	Recovery .		Recovery.	до до	Death	Recovery.	ор	do	doRecovery.	ор	do
	Remedy.	None	Potassium permanga- nate.	Potassium perman an ganate (put into ru-	1 1 4	manganate. Caffein Potassium per-	manganate. Charcoal; atro-	Charcoal	AtropinTannic acid	doses). None Godium bicar- e ar b on ate (represented)	doses). Tannic acid (repeated	doses). Nonedo
	Severity of illness.	Sick	Not sick	Slightly sick	Symptoms Slightly sick Sick	Death Sick	Very sick	do	do. Slightly sick.	Very slek Not siek Very siek	Sick	Very sick
Part of plant used (fed	unless otherwise stated).	Seed heads, fully de-	Seed heads, fully developed, mixed with potassium perman-	ganate (drench). Seed heads, fully developed (forced feeding).	do. do.	op.	ор	Seed heads, ripening	Seed heads, fully developed (forced	feeding). do. do.	do	do
	Date of experiment.	July 11	do	July 13	.do July 14.	July 15.	do	July 22.	July 23.	Aug. 10. Aug. 11. Aug. 13.	Aug. 14	.578 Aug. 15
Woightof	plant used.	Pounds. 0.348	.340	. 392	. 463 . 683 . 899	.741	. 736	888.	.838	. 811 . 612 . 939	. 886	.578
	Weight.	Pounds. 79 to 74	77 to 76	89 to 85	105 to 98 90 to 88 102 to 96	84 81½ to 70	83½ to 85	120 to 108	113 to 100 91 to 88½	92 to 80½ 92 106½ to 101	100½ to 91	65½ to 60½
Animal.	Designation.	I.—Sheep given Z. venenosus—Con. No. 234.	No. 253	*No. 252.	*No. 264 No. 254 No. 268	No. 262 No. 283	No. 290.	No. 259	*No. 247	*No. 269. *No. 231 *No. 259.	No. 267	*No 294

Do.	Do.	До.	Do. Do.	Do.	Do. Do.	Do. Do. Do.	Fishlake National Forest, Utah.	Do. Do.	Ephraim, Utah. Do.
.881		.882		1, 102	. 876 . 919 . 540	.110 .219 .197 .220	219 Fish	. 447 1. 322 . 882 1. 689	.551 Eph
	1.102	ω. 	H.H.	Ξ	w, w, a,			7.1.3.1	25, 4,
do	dodo	do	op.	· Death	Recovery do do do	do Recovery. Deathdo			Recoverydo
Tannic acid	Magnesium Sulphate; tannic acid (repeated doses); atro-	Sodium bicar- bonate (re- p e a t e d	dodo	Sodium bicar- bonate (re- p e a t e d	doses). dodoBled	Nonedo Bleddo	None	do do do	op.
Siek.	doVery sick	Sick.	Very sick Not sick	Death	Siek. Very siek. do.	Slightly sick. Death. Very sick. Death.	Not sick	Not sick (?) do. Not sick	Slightly sickSymptoms
do.	do.	do.	do. Seed heads, fully developed (in bran).	Seed heads, fully developed (forced feed- ing).	Seed pods, with seed removed (forced	Seed (forced feeding) do do	Leaves, stem, flowers, and pods (forced	Leaves, stems, flowers, and some young roads (formed feeding)	Leaves, stems, and fruit (forced feeding).
do.	Aug. 17.	Aug. 21.	Aug. 22. Aug. 24. Aug. 25.	op	Aug. 26	Sept. 11 Sept. 12 Sept. 12 Gept. 13	Aug. 8	Aug. 9. Aug. 11.	June 10do.
.639	. 981	. 600	.959 1,506 .368	1.047	. 705	. 087 . 145 . 149 . 152 . 153	. 244	. 434 . 846 . 723 1. 267	. 444
72½ to 72½	89 to 843 86 to 772	68 to 61	87 to 77½ 102½ 111 to 106½	99	80½ to 81 84 to 68 77 to 68	79 66 77 76 69	111	97 64 to 60 82 to 78 75 to 713	803
*No. 296	*No. 265 *No. 291	*No. 293	*No. 292 *No. 246 *No. 247	*No. 264	*No. 240 *No. 277 *No. 299	No. 235. No. 251. No. 245. No. 297. No. 266.	No. 233	No. 246 No. 281 No. 292 No. 232	Z.—Sheep given Z. paniculatus: No. 278.

Dranch

Following are the details of three cases which may be considered typical:

Sheep No. 193.—This animal (Table I, section G) was a 2-year-old ewe, lent for experimental purposes by Mr. Ole Birkeland. She was received at the station on May 9, 1913. An attempt was made on May 12 to feed to her the bulbs of Zygadenus venenosus ground and mixed with bran. As she would not eat this, a trial was made on May 13 of feeding her with Zygadenus tops, but these also she refused to eat; so at 1.40 p. m. of the same day she was drenched with 200 grams of Z. venenosus bulbs ground fine and suspended in water. At 2.35 p. m. she was frothing at the mouth and vomiting, with violent contractions of the diaphragm and abdominal muscles. She was lying down, but was able to stand. At 3.50 p. m. she was still frothing at the mouth, but was fairly strong. At 5.30 p. m. her temperature was 101° F. She was fairly strong, but acted as though in much discomfort. On May 14, at 7 a. m. she appeared entirely normal, and at 8.30 she was turned out to pasture.

On June 14, 1913, she was kept in for feeding, and at 8.15 a. m. she was fed 100 grams of fresh tops of *Zygadenus venenosus*, which were collected on June 12. At this time the plant was in flower. At 4.40 p. m. she was fed 200 grams of the plant, and at 7.40 p. m. 135 grams.

On June 16, at 7.10 a.m., she was fed 200 grams; at 6.40 p.m., 200 grams; and at 7.15 p.m., 250 grams. All the Zygadenus fed on June 16 had been collected on the preceding day.

On June 17, at 6.45 a. m., all the Zygadenus given on the preceding day had been eaten. The animal appeared bright, but showed sensitiveness to sudden noises and there was some trembling of the surface muscles. At 7 a. m. she was run around the corral. It was found that she moved in a stiff-legged manner and was somewhat weak. The stiffness was most noticeable in the hind legs. She was licking her lips and rubbing her nose against the fence and moving her head about in a jerky way. At 8.45 a. m. the symptoms were about the same as at 7. a. m. At this time she was fed 200 grams of Zygadenus. At 11.30 a. m. and 1.50 p. m. she was fed 100 grams of Zygadenus. At 3 p. m. her temperature was 103.8° F. She was more nervous than in the morning and her movements were somewhat more stiff. The jerking movements were more pronounced and continuous. All the Zygadenus which had been previously given had been eaten. At 8 p. m. the symptoms as noted at 3 o'clock still continued. She was fed a little alfalfa.

On June 18, at 6.45 a.m., the sheep was trembling almost constantly, with frequent spasmodic movements. Her legs were stiff as she walked about, and there was some lack of control. She was dejected and dull and without appetite. At 9 a.m. she was given 5 grams of tannic acid. At 9.40 she was given 4 ounces of Epsom salts in solution. At 10.30 a.m. she was put in a metabolism cage, in order to collect the excretions. At 2 p.m. she seemed to be decidedly better, and continued in about the same condition throughout the afternoon.

On June 19, at 6.30 a. m., the animal trembled, but showed no other symptoms. She had not urinated since being placed in the metabolism cage at 10.30 a. m. on the preceding day. At 8 a. m. she was taken out of the cage and fed some alfalfa. The general appearance of the animal was better than on the preceding day, but she was still unsteady in her gait and exhibited trembling of the surface muscles. At 1 p. m. she defecated as the result of the dose of magnesium sulphate given on the preceding day, and this defecation continued in a mild diarrhea. At 3 p. m. she urinated for the first time after being placed in the metabolism cage. At this time her respiration was 148, and she seemed in general to be worse. At 8 p. m. she was given 1 gram of diuretin in solution, and returned to the metabolism cage. During the afternoon the animal seemed to be gradually getting worse. When standing, her hind legs were

drawn forward under her. The muscles of the legs were twitching almost continuously, and it was with difficulty that she could get up and down.

On June 20, at 6.30 a.m., there were about $1\frac{1}{2}$ pints of urine which had accumulated through the night. This was preserved in alcohol, and a chemical examination showed that it contained the alkaloids of Zygadenus. The diarrhea still continued. The general condition of the animal was nearly the same as the preceding night, except that she appeared a little stronger. She was taken out of the cage and fed alfalfa. At 2 p. m. she was somewhat better than in the morning and had an appetite, although she still preferred to lie down. At this time she was given 1 gram of diuretin in solution. At 7.50 p. m. she was much better, standing in a more normal manner and with no noticeable trembling. On June 22 she was turned out, apparently all right.

On July 15, 1913, the sheep was again kept in for feeding, and on July 16, at 10.30 a.m., she was given 200 grams of the mature heads of $Zygadenus\ venenosus$, consisting of

pods and seeds, ground and mixed in bran.

On July 17, at 9.25 a. m., she was fed 270 grams prepared as the day before, and at 1.25 p. m. she was fed 180 grams. At 7.45 p. m. she was fed 210 grams. At the time of the last feeding she showed the effects of the poisoning. She did not move with the usual freedom, and there was some twitching of the surface muscles of the body.

On July 18, at 9.50 a. m., she was fed 235 grams. At 6.30 p. m. she was fed 220 grams. During the day there was little change in the condition of the animal.

On July 19, at 7 a. m., the symptoms were much more pronounced than the preceding night. There was stiffness of movement of the legs, licking of the lips, and slight trembling. This stiffness and accompanying clumsiness were more pronounced in the hind legs. At 9 a. m. she was fed 255 grams and at 3 p. m. 195 grams. At this time the animal was considerably weaker than in the morning.

On July 20, at 8.45 a.m., all the Zygadenus given on the preceding day had been eaten, and the general condition of the animal was about the same as on the preceding night. She moved with some difficulty and with marked stiffness of the legs. There was trembling of the surface muscles accompanied by some shaking of the head and licking of the lips. She was unusually sensitive to noise, as she was easily startled, and at such times there were sudden contractions of the body muscles. She had much difficulty in raising her feet sufficiently to get over elevations of 3 or 4 inches.

On July 21, at 3 p. m., the animal had improved in its general condition, although the symptoms were still well marked. These did not differ from those seen earlier in

the sickness, but were less pronounced.

On July 22, at 7 a. m., the animal moved about fairly well, but there was some trembling of the surface muscles, especially in the shoulders. She improved during the day and at 7 p. m. seemed to be quite normal. On July 23, at 7.55 a. m., she was turned out to pasture, appearing strong and active and showing no symptoms except some slight trembling when handled.

Sheep No. 160.—This animal (Table I, section F) was a ewe lent by Mr. Ole Birkeland on June 20, 1912. On July 19, 1912, she was taken in for feeding with Zygadenus venenosus. At 11.05 a.m. her respiration was 28, pulse 26, and temperature 105° F. At 11.25 a.m. she was drenched with 125 grams of the seed heads of Zygadenus venenosus suspended in water. These seed heads included the pods and seeds. At 11.35 a.m. there was a little frothing at the mouth. At 11.40 a.m. her respiration was very rapid and irregular. It would run as high as 200 per minute for perhaps 50 respirations, then stop, only to be resumed at the same rapid rate. At this time the animal had vomited. At 12.55 her pulse was 95 and very variable; her temperature was 104° F. At 1 p. m. her respiration was 200 or more and the heart action regular and strong. She was then given a solution of potassium permanganate. At 1.17 p. m. she was getting weaker on her legs. She would start to lie down and nearly fall, but would get on her feet after a minute or two and then lie down again. Her respiration was about 180 per minute. At 1.30 p. m. her respiration was still rapid, the mucous

membranes of the mouth somewhat cyanotic, and the ears drooped. At 1.47 p. m. she was given subcutaneously 8 grams of caffein. At 1.49 p. m. she threw her head up and held her breath for about a quarter of a minute. She repeated this action several times. At 2.03 p. m., her temperature was 100° F. The animal seemed somewhat stronger. Her respiration was still variable, running as high as 180 per minute. Her pulse was about 80. She was still frothing at the mouth and appeared to be in pain. At 2.20 p. m. she had a hard time to breathe. She shook her head, staggered about, and lay down. Her respiration was slow and labored. At 2.22 p. m. her respiration was getting more rapid. At 2.23 p. m. the animal seemed to be in great pain; her breathing nearly stopped for a minute, but started again in a panting manner. Her respiration continued irregular, first fast and then slow and labored. At 3 p. m. her temperature was 100.2° F. and her respiration 84, but not as labored as at 2.03 p. m. She was still frothing at the mouth. Her pulse was 96 and strong. At 3.30 p. m. her respiration was 160. She fell upon her knees, struggled to get on her feet again, but finally lay down. At 4.15 p. m. she was groaning more or less and staggered as she attempted to walk. Her respiration was 90. At 4.30 p. m. she was lying down and her respiration was very nearly normal. Her temperature was 99.1° F. At 5 p. m. she was down again and in pain, groaning continuously, her respiration slower than for some time previous, the rate being about 60 per minute. At 5.04 p.m. the animal was lying sprawled out upon her belly. Her respiration was 168. 5.38 p. m. she appeared very stupid, almost as if sleepy. She was given subcutaneously 6 grains of caffein sodiobenzoate. At 5.45 p. m. her temperature was 99.4° F., her respiration 120 and very variable, and her pulse 96. At 7.10 p. m. her temperature was 99.7° F. Her respiration was slow and variable. After expiration there would be a pause followed by two short and shallow inspirations close together. These would be followed by a long inspiration, then a full expiration accompanied by a groan. This was repeated over and over again, the whole cycle taking about 20 seconds. The animal on the whole seemed to be brighter than at 6 p. m.

At 8 p. m. the animal was given 5 grains of caffein sodiobenzoate. Her temperature was 100.8° F., respiration 10, and pulse 120. At 10 p. m. her temperature was 100.2° F., pulse 120, and respiration 4. The inspiration was deep and the expiration was accompanied by a groan. At 10.55 p. m. her temperature was 100° F. At 11.15 p. m. her respiration was 18 and pulse 108. Her general condition was unchanged.

On July 20, at 7 a. m., the animal's pulse was 100, temperature 100.8° F., and respiration 11. She was then lying with her head bent under her body and would probably have died in that position had she not been relieved. She seemed at this time unconscious. At 8.45 a. m. her temperature was 100.6° F, pulse 84, and respiration 12. At 10 a. m. her respiration was 10, and her pulse 84. The animal was in a comatose condition. At 10.50 a. m. her respiration seemed to be getting more shallow. At 11.15 a. m. she seemed somewhat brighter than earlier in the day. At 12 m. her temperature was 102.6° F., respiration 12, and pulse 108. During the afternoon she had been lying in practically the same position, with her head slightly raised, resting upon a support. She was too weak to move herself at all. At 3 p. m. her respiration was 12. At 3.30 she was given subcutaneously 10 c. c. of whisky. At 4.20 p. m. her temperature was 102.8° F., respiration 36, and pulse 116. At 8.05 p. m. she was given subcutaneously 5 c. c. of whisky. At 9.30 p. m. her temperature was 104.6° F., respiration 18, and pulse 120.

On July 21, at 5.45 a. m., she was found in practically the same condition as the preceding night. At 6.45 a. m. her temperature was 104° F., respiration 24, and pulse 148, and weak. At 9.50 a. m. she was given subcutaneously 5 drops of fluid extract of digitalis in 8 c. c. of whisky. At 10.15 a. m. her pulse was somewhat stronger than before the digitalis was given. At 11 a. m. her pulse was 102, temperature 104.6° F., and respiration 48. At 11.15 a. m. her respiration was fairly deep, but was somewhat

spasmodic. Her pulse was weak. At 11.45 a.m. her respiration was 68. At 9.30 p.m. her temperature was 105.3° F., respiration 24, and pulse 120, but weak. The animal was given subcutaneously 3 drops of fluid extract of digitalis in 6 c. c. of whisky. During the day there had been very little change in her condition. She lay in a coma, from which she did not rouse herself except occasionally to shake the flies from her ears. Her position had been changed from time to time by the attendants. She was found dead on the morning of July 22.

At the autopsy the venous blood vessels were found congested and the lungs were congested, as were the liver and kidneys. There was considerable inflammation of the walls of the fourth stomach and of the whole length of the intestines. Sections of the kidney showed that the capillaries were much congested, and there was some degeneration of the tubule walls. Sections of the liver showed acute congestion, and the same condition was noticed in the sections of the lung.

Sheep No. 197.—This animal (Table I, section G) was a ewe 2 years old, lent by Mr. Ole Birkeland on May 9, 1913. An unsuccessful attempt was made to feed Zygadenus venenosus tops to her on May 12.

On May 26, at 11.25 a. m., she was drenched with 178 grams of Zygadenus venenosus tops suspended in water. These plants were collected on May 23. At 11.40 a. m. she was frothing at the mouth. At 11.45 a. m. she was given a drench of 1 gram of diuretin and 0.455 gram of caffein citrate. At 11.50 a.m. she was vomiting, and when observed at 12 m. the vomiting was continuing. At 12.05 p. m. her respiration was getting irregular and deeper. At 12.30 p. m. her respiration was extremely fast and she was panting. At this time she was violently nauseated and threw herself down two or three times and then jerked about in a spasmodic manner. At 12.40 p. m., being extremely nauseated she was trying to vomit, throwing herself down, and the spasmodic movements were followed by quick, panting respiration. At 12.45 p. m. her respiration was about 200. She showed weakness in her legs. At 1.15 p. m. she repeated the spasmodic movements which had been noticed at 12.30 and 12.40 p. m., evidently struggling to get breath. She threw her head from side to side and ran the length of the corral, throwing herself upon the ground and rising again as though having a fit. The mucous membranes of the mouth were cyanotic. These movements were repeated a little later. At 1.30 p. m. she was given a dose of 5 c. c. of gin. Another struggle for breath followed, and it was noted after this struggle that her heart action was very rapid and strong. The beat was audible to the observers. Five c. c. more of gin were given subcutaneously. At 1.36 p. m. she was lying upon her side. Her respiration was 160. At 1.40 p. m. she had another struggle for breath, throwing herself about violently, even throwing herself over upon her back. These struggles were repeated at 1.49 and at 1.53 p. m. The mucous membrane of the mouth at both times was very markedly cyanotic. At 1.55 p. m. she was given 5 c. c. of gin. At this time she was still strong enough to get on her feet. She was urinating freely. At 1.59 p. m. she passed through another spasmodic attempt to breathe. At 2.02 p. m. her pulse was about 200. At 2.06 p. m. there was a spasmodic struggle for breath. At 2.10 p. m. the animal was breathing with very great difficulty. Ammonia was used to stimulate her respiration. Her pulse was 130. At 2.45 p. m. she had great difficulty in respiration, but at this time it was not accompanied by a spasmodic struggle. At 3.06 p. m. she had another spasmodic struggle, and ammonia was used as a stimulant. At 3.30 p. m. her respiration some of the time was very rapid, becoming as high as 200 per minute. Then it slowed down and became labored. On the whole the animal at this time seemed somewhat better. At 3.39 p. m. she made a struggle to get upon her feet but was unable to do so. Her respiration at this time was variable and very labored, the breathing being followed by quick, panting efforts. At 4.10 p. m. her respiration nearly stopped. She was stimulated with ammonia. At 4.15 p. m. ammonia was again used. At 4.45 p. m. her respiration was 132. At

5.19 p. m. the animal was upon her side. Her respiration was labored, but seemed somewhat stronger. Her condition remained very nearly the same until 6 p. m. At 6.30 p. m. she was found dead.

The autopsy showed that the lungs were slightly congested; the inner walls of the ileum were congested and the venous blood vessels, generally speaking, were full. It was evident that death had occurred from respiratory paralysis.

The detailed report of the examination of the internal organs by Dr. Mohler, of the Bureau of Animal Industry, is as follows:

Kidney (cortex and medulla).—Many of the intertubular capillaries in the labyrinth of the cortex and some Malpighian bodies show a marked distention, but not sufficiently pronounced to be called congestion. There is also a general distention of many of the convoluted tubules and the interstices between the capsule of Bowman and glomeruli with a serous, cedematous exudate. This latter has distended the tubules and compressed the renal epithelial cells, many of which, having become atrophic from pressure, disintegrated and desquamated into the lumen of the tube. While these changes are quite marked in the convoluted tubules, the cedema, desquamation, and degeneration are absent in the straight portions of the uriniferous tubules of the medulla. The distention of the capillaries, however, is present even in the medulla. No interstitial alterations are present.

Lung.—The characteristic lesion is the intense congestion of the entire organ, the presence of small lobular areas of consolidation, and occasional minute cedematous areas. The larger pulmonary and bronchial vessels are all overdistended, but the interfundibular capillaries show not only overdistention but also diapedesis and outwandering of the leucocytes. No such capillary hemorrhages or poollike accumulations of the blood can be seen in this lung as were previously observed in lung 716. The bronchial tubes and the smaller bronchi are unaltered. There is no peribronchitis present, although the bronchial blood vessels have all participated in the distention of the other vessels of the lung.

Kidney.—No acute inflammatory changes present. Evidences of a slight subacute catarrhal nephritis accompanied by mild degenerative changes in the renal cells in the cortical portion of the kidney. No interstitial changes present.

Liver.—Moderate amount of physiological fatty infiltration and a slight congestion of the intralobular capillaries between the liver cords, but no diapedesis of red blood cells or outwandering of leucocytes. The hepatic cells proper show a slight amount of cloudy swelling in isolated lobules. No interstitial changes present.

Ileum.—Shows a slight increase in the adenoid tissue in the mucosa and slight fullness of the blood vessels in the submucosa. There is, however, no congestion, desquamation, or degeneration present.

SYMPTOMS IN SHEEP OBSERVED AT THE GREYCLIFF STATION.

The very large number of cases of illness and death observed at the Greycliff station furnished an opportunity for a fairly complete picture of the symptoms produced by Zygadenus poisoning. The symptoms were noted in detail, and the description that follows is drawn from a summarized statement formulated from these notes.

SALIVATION.

Generally salivation, or frothing at the mouth, was the first noticeable symptom and continued through the acute period of the illness. It was not invariably present; sometimes it did not appear, especially in the fed cases. It was seen in nearly all the drenched

cases, and was rarely absent when the attack was acute. The salivation was in many cases accompanied by grinding of the teeth. Plate IV, figure 1, of sheep No. 160, and Plate IV, figure 2, of sheep No. 192, illustrate this stage of the illness.

NAUSEA.

Nausea was very pronounced in nearly all cases, and frequently resulted in violent vomiting, this vomiting, like the salivation, being largely confined to the acute stage of the illness.

PULSE.

Routine observations upon the pulse were made in a large number of cases. The rate of the pulse is, of course, very variable under normal conditions. When taken before the experimental feeding it varied from 52 to 144, although in most cases it was between 60 and 100. Generally speaking, when the intoxication was not acute there was very little change from what would be expected in normal variations either in the rate or character of the pulse. In the severer cases, especially in those that ended fatally, the rate was from 125 to 200. While in three cases of sheep not under the influence of a toxic substance the pulse was 144, this condition is unusual; and in a general way it seems to be true that if the rate runs much above 130 a fatal termination of the illness is likely to follow. In the severe cases the pulse was weak and sometimes intermittent.

TEMPERATURE.

Temperature observations were made in detail in a large number of cases. It was considered necessary to get the average of a considerable number, inasmuch as there is in sheep quite a range of variability under normal conditions and also a considerable difference in individuals. The extreme range of temperature was from 97.4° to 105.7° F. From the cases of 1914, 64 records were made. Of these, 8 showed no marked change, 14 exhibited an increase, and 42 a decrease, and the decrease ordinarily was not very great but in some few cases was down to between 97° and 98° F. It is evident that, in general, intoxication by Zygadenus is accompanied by depression of temperature. In some few cases, in which there were no other symptoms of poisoning, a lowering of temperature was noticed; this, however, was not sufficiently general so that it could be considered diagnostic in the absence of other symptoms. Curves are given (figs. 1 and 2) of sheep 282 and 291. These, it should be stated, are not average cases, but they may be considered typical of cases in which the lowering of temperature is more marked.

RESPIRATION.

The rate of respiration had an extremely wide range of variation. Quite uniformly in the acute stages of the poisoning, the rate was very rapid, running in some cases as high as 250 per minute. After this period the rate was very much reduced, falling to normal or below,

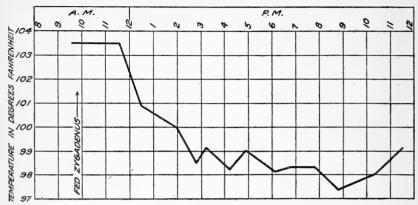


Fig. 1.—Curve of temperature of sheep No. 282.

and the animal sometimes lay for hours breathing most of the time in a slow and labored fashion. This period of comparative quiet might be interrupted, sometimes frequently, by times of rapid breathing, accompanied by panting and followed quickly by a very slow rate. Sometimes, in severe cases, there were times when the animal threw

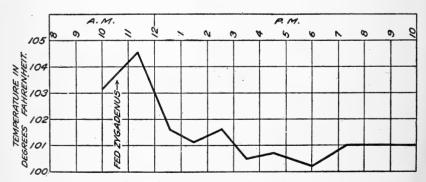


Fig. 2.—Curve of temperature of sheep No. 291.

itself about violently, fighting for oxygen. This condition lasted for perhaps two or three minutes and was succeeded by a period of quiet, which was soon broken by another struggle. During these struggles the mucous membranes of the mouth were frequently cyanotic. The struggles were spasmodic, and when authors state that poisoned animals have spasms or convulsions, it is to be presumed that they

refer to this condition. It should be noted, however, that in the cases observed at the Greycliff station there was no indication of any special tonic or clonic contraction of the muscles; the violent movements of the animals were simply those caused by distress from dyspnæa.

Figure 3 gives the curve of respiration for sheep No. 174 and may be considered typical of the average fatal case. The sheep was

drenched at 12 o'clock noon and died at 11.15 p.m. Therespiratory rate rose to 200 between 2 and 3 o'clock, when the animal had one of the spasmodic struggles for breath. It then fell to 9 and remained low, with comparatively slight variations, until the time of death.

Figure 4 gives the curve of respiration of sheep No. 160, a prolonged case. This animal was drenched with Zygadenus at 11.25 a. m., July 19, and died during the night of July 21. respiration almost immediately after the dose was given ran up to 200 and during the afternoon varied between 60 and 168.

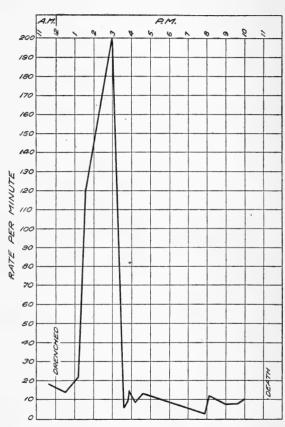


Fig. 3.—Curve of respiration of sheep No. 174.

In the evening it fell, and after that time the maximum noticed was 68, but most of the time it was near 20 or 30.

MUSCULAR WEAKNESS.

In all cases of any severity muscular weakness was noticeable. Early in the illness the animals staggered, and in the more serious cases not only could not rise, but lay flat upon the ground. This weakness was most pronounced in the forelegs. Plate V, figure 1, shows this condition of weakness in the forelegs in sheep No. 162, while Plate V, figure 2, shows the same animal down. Plate VI,

figure 1, shows sheep No. 174 when down and very sick. This picture was taken just before a spasmodic struggle for breath.

In many cases in which the animals were strong enough to remain on their feet, the gait was peculiarly stiff legged. Both fore and hind limbs were affected, but the condition was most pronounced in the hind legs. Sometimes the hind legs were moved less readily, approximating, perhaps, the condition noticed by other writers in laboratory experiments. These symptoms were especially noticeable in the cases in which the animals were fed and were not very sick.

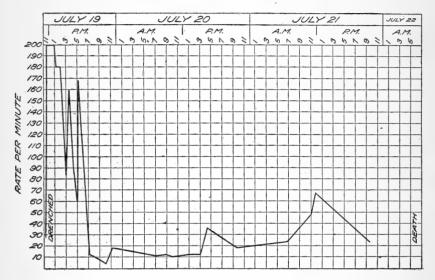


Fig. 4.—Curve of respiration of sheep No. 160.

TREMBLING AND HEIGHTENED REFLEXES.

The fed cases generally exhibited trembling and a sensitiveness to sudden noises or movements. A blow upon the corral fence was followed by a sudden start on the part of the animal, or a light blow upon the animal was followed by a quick reflex movement. This condition was not noticed in the drenched cases and seems to be more characteristic of prolonged illnesses. As stated before, this symptom of heightened reflexes had been noted by both Chesnut and Hunt.

COMA.

While, as already noted under the head of respiration, death resulted from respiratory failure and was frequently preceded by spasmodic attacks of dyspnœa, there were other cases in which the animals lay quietly hour after hour, and sometimes even for days, with labored breathing, in a condition of coma which ended in



FIG. 1.—SHEEP No. 168 AT 1.30 P. M., SHOWING WEAKNESS IN FORELEGS.

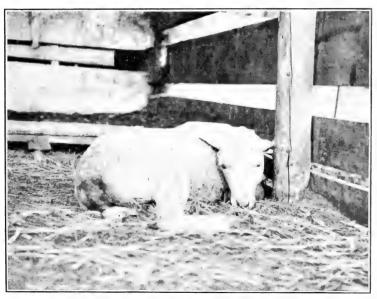


FIG. 2.—SHEEP NO. 168 AT 5.45 P. M., WHEN UNABLE TO RISE.



Fig. 1.—Sheep No. 174, Down and in Bad Condition. Photographed Just Before a Spasmodic Struggle for Breath.



Fig. 2.—Sheep No. 161, Down Almost Two Hours and Unable to Rise.

death without any exhibition of spasms. Plate VI, figure 2, shows sheep No. 161 in this condition of coma. Sheep that are poisoned on the range are more apt to be in this condition of prolonged coma than to show the more violent symptoms of dyspnæa exhibited by animals that are drenched or forcibly fed.

SYMPTOMS IN HORSES AND CATTLE.

No results on horses were reached in the experimental work at Greycliff. From conversations with stockmen who have had experience with horses poisoned by Zygadenus, it appears that, in general, the symptoms resemble those exhibited in sheep.

In the cases of the two head of cattle which showed symptoms, the experiment was carried only to the point of proving the toxic effect of the plant, and no attempt was made to get a complete symptomatic picture. The animals became uneasy, displayed heightened reflexes, and one dragged the hind legs slightly. So far as they went, the symptoms were like those observed in sheep.

AUTOPSIES.

Autopsies were made on four cases in 1912, six in 1913, and seven in 1914. Of the cases in 1913, sheep No. 186, while showing distinct symptoms of Zygadenus poisoning, died as the result of the administration of morphin.

The appearances presented by these animals at the autopsies were quite uniform, though not alike in all details. In six cases there was epicarditis. In nearly all, the inner wall of the ileum was hyperæmic or congested, and in all but one the lungs were congested. The kidneys were congested and more or less degenerated in most cases. Generally the heart was in systole, the contraction being most marked in the left ventricle.

Generally speaking, then, the post-mortem appearances may be stated as including inflammation of the inner wall of the ileum and occasionally of the fourth stomach and large intestines, the heart in systole, congestion of the lungs, and congestion and more or less degeneration of the kidneys. Possibly the condition of epicarditis may be considered typical, although it was not noted in all the cases.

The preserved material from the autopsies was examined by Dr. Mohler, and the following summarized statement of the pathological findings and the inferences to be drawn has been furnished by him:

The most conspicuous phenomenon shown in all six cases was the high capillary blood pressure, manifested principally in those organs which eliminated the active principle of the ingested substance, that is, kidneys, lungs, and liver, being also shown to a less extent in the intestine.

In the kidney the changes in the capillary varix were fullness to overdistention, which was accompanied by outwandering of leucocytes, diapedesis of the red blood

corpuscles, and all the phenomena of a congestion or an acute or subacute inflammation. Occasional ruptures of the capillary vessels were noted, forming poollike capillary hemorrhages.

The distention of the capillaries in some instances had brought about cloudy swelling, or the early stage of degeneration of the renal epithelium, which in some instances had become desquamated. The supporting or interstitial tissue was not affected.

In the lung the high capillary pressure is even more manifest than in the kidney, owing to the presence of a greater number of capillaries. The variations were from moderate fullness to overdistention, followed by inflammation in the more acute cases, resulting in localized areas of ædema where the serum had oozed out and filled one or more lobules of the lung.

In spite of the fact that the fullness, congestion, and inflammation were more marked in the lung than in the kidneys, the degenerative changes and the desquamation of the pulmonary epithelium were less evident and not as frequent, owing to the greater resistance of the pulmonary cells. While no interstitial changes were present in the kidney, slight interstitial changes in the lungs were present in the peribronchial areas in some of the cases. In others, the interstitial changes were also present in the visceral pleura.

In the liver the vascular changes were either entirely absent or so slight as not to deserve any mention, but the epithelial changes were quite marked, owing to the more delicate composition of the cytoplasm. The absence of vascular changes indicates that the elimination by this organ is but very slight and that the metabolic function is quite able to take care of any of the irritant products that may have reached the liver.

In the intestine the vascular changes are likewise very slight.

TOXIC AND LETHAL DOSE OF ZYGADENUS VENENOSUS FOR SHEEP.

The very large number of feeding experiments with sheep at Greycliff made it possible to determine the toxic and lethal dose with considerable accuracy. Inasmuch as very little has been known in regard to the toxic dose of Zygadenus for sheep, the results of these cases are especially interesting. Table II summarizes the nonfatal cases, showing their number and the quantities of the plant necessary to produce illness.

Table II.—Nonfatal cases of poisoning of sheep by Zygadenus venenosus at Greycliff, Mont., in 1912, 1913, and 1914.

Feeding experiments.	Number of cases.	Quantity used per 100 pounds of animal.		
		Maximum.	Minimum.	Average.
Season of 1912: Drenched with leaves, stems, and fruit. Drenched with stems and leaves. Drenched with stems, fruit, and some leaves. Drenched with fruit. Season of 1913: Fed on leaves. Drenched with leaves. Drenched with bulbs. Drenched with leaves and buds. Drenched with leaves and flowers. Fed on leaves and flowers. Fed on leaves, flowers, and fruit. Fed on seeds. Fed on seeds.	2 1 2 5 4 8 8 3 1 1 1 6 2 2 2 1 3	Pounds. 1. 324 2.1 264 2.1 747 612 4.6 496 7.188 5.597 241	Pounds. 0, 79 33 141 893 385 517 1, 728 495 3, 155	Pounds. 1. 057 . 33 . 55 . 228 1. 607 . 5746 . 575 . 389 2. 7513 . 4955 5. 1715 5. 597 . 1613

Table II.—Nonfatal cases of poisoning of sheep by Zygadenus venenosus at Greycliff,
Mont., in 1912, 1913, and 1914—Continued.

Feeding experiments.	Number of cases.	Quantity used per 100 pounds of animal.		
		Maximum.	Minimum.	Average.
Season of 1914: Forced feeding of leaves and some young buds. Forced feeding of leaves and some young buds. Fed on leaves, some young buds, and a few flowers. Forced feeding of leaves, stems, flowers, and buds— Material collected near the station. Cabin Corral collections Fed on leaves, stems, flowers, and buds. Forced feeding of leaves, stems, and flowers (Cabin Corral collections). Forced feeding of leaves, stems, flowers, and young fruit. Forced feeding of very young seed heads. Forced feeding of seed heads, some fully developed and others half developed. Torral collections). Forced feeding of half-developed seed heads (Cabin Corral collections). Forced feeding of nearly developed seed heads. Forced feeding of fully developed seed heads. Forced feeding of ripening seed heads. Forced feeding of ripening seed heads. Forced feeding of pods with seeds removed. Forced feeding of seeds.	4 1 9 4 1 1 3 6 18	Pounds. 0.662 1.983 2.756 .992 1.754 1.323 .994 .741	1. 436 .992 .757 1. 543 .881 .440 .740	Pounds. 0.6615 .5245 1.643 .5495 1.789 1.912 1.712 .851 1.543 1.432 1.69 1.175 .859 .7415 .540 .153

Table III summarizes the fatal cases in the three seasons.

Table III.—Fatal cases of poisoning of sheep by Zygadenus venenosus at Greycliff, Mont., 1912, 1913, and 1914.

Feeding experiments.	Number of cases.	Quantity used per 100 pounds of animal.		
		Maximum.	Minimum.	Average.
Season of 1912: Drenched with fruit Season of 1913: Fed on leaves Drenched with leaves Drenched with leaves Drenched with leaves and flowers Season of 1914: Forced feeding of leaves, stems, and flowers Forced feeding of fully developed seed heads Forced feeding of seeds.	1	Pounds. 0.853 .746	Pounds. 0, 299 .384 .882 .199	Pounds. 0.571 1.397 .537 .550 .544 .936 .213

As these feedings were carried on during the season as long as the plants could be obtained and as it was practically impossible to have any considerable number of cases at one time, it is evident that the number of cases under any given set of conditions must have been small. As a matter of fact, none of the cases of 1912 were strictly comparable with those of 1913. Consequently, the actual averages of dosage were based on a comparatively small number of cases.

In 1914, there was a much larger number of cases, and some stages of the plant were fed upon which no experiments were made in the preceding years. Even in this year, however, there were only a few cases in which the experiments were under identical conditions.

In the compilation of Tables II and III some of the cases have been excluded. In Table III all cases in which the remedy given was clearly effective were excluded, for some of these received what would have been a lethal dose had it not been for the remedy. In Table III cases were excluded which were known to have received much more than a lethal dose. The uniformity of dosage in 1914 is explained by the fact that the preceding work had shown clearly that the toxic dose was not far from 0.5 pound, and the experiments were made on this basis. It should be noted, too, that most of the work of the summer of 1914 was with reference to the experimental use of remedies, so that the quantity of the plant administered was estimated to be sufficient not simply to produce symptoms, but to make the animal very sick, in order to get a fair test of the remedy. Hence, the average figures for the toxic dose will be rather high.

The "forced feeding" cases of 1914 can be fairly compared with the "drenched" cases of 1912 and 1913, as the difference between the two methods is mainly in the fact that in forced feeding no water is used while in drenching considerable water is necessary as a vehicle for the weed.

The age of the animal played a comparatively small part in these experiments, as all the animals were mature, most of them being 2 years old or older.

It will be noticed that when the plant was given in the form of a drench or by forced feeding, the dosage, as would be expected, was considerably less than when it was given with food. An examination of the complete table of feeding (Table I) shows also very clearly that the size of the dose varied inversely with the time during which the material was eaten. In a large number of cases in which the plant was given with food, the feeding extended over two or more days. In those cases the dosage was considerably greater than when the material was fed in a single day. It may be assumed that if the same quantity of the plant which was received in a drench could have been fed within a short period of time, the effect would have been the same.

The average dose which produced illness when administered in the form of a drench or by forced feeding was practically the same for all parts of the plant except the pods and seed. It appears that the plants are less toxic at the time when the pods are forming, which may be due, in part at least, to the diminished toxicity of the leaves as they dry up. It is not clear, however, from this work, that the leaves lose any appreciable amount of toxicity, and the more probable explanation is that the pods at this time are only slightly toxic. In the single experiment of feeding pods without seeds, the dosage was about like that of other parts of the plants, but it is probable from the detailed history of this experiment that this is not a fair representative of such cases.

The seeds are very much more poisonous than any other part of the plant. Heyl, Loy, Knight, and Prien (1912, p. 17) give the results of determinations of alkaloids in different parts of the plant. Their statement is obscure and contradictory, but apparently they reach the conclusion that the bulbs and leaves contain approximately the same quantity of the alkaloid, the roots much less, and the flowers about twice as much as the bulbs and leaves. This compares very well with the results of the experimental feeding at Greycliff, except that it did not appear that the flowers were more toxic than other parts of the plant.

Table II gives the maximum and minimum dosage, and it will be noticed that there is a considerable range of variation between these two. The individual peculiarities of the animal in cases of poisoning doubtless must be taken into account, and the detailed table of the experiments shows that in some cases a larger quantity of plant than that which this table indicates to be toxic may be administered without effect. In most of the cases, however, where the larger amount was used, the feeding was distributed over a

longer time.

In general, the experiments seem to indicate that when any part of the plant except the seed is used the toxic dose varies from 1.6 pounds per hundredweight of animal to 5.6 pounds, this wide range of variation being accounted for by the more or less extended time of feeding. In the drenching and forced-feeding experiments, more uniform results were reached, showing that the toxic dose of all parts of the plant, except the seed, is not far from 0.5 pound per hundredweight of animal.

There is considerable difference in the items of the summarized tables in the exactness of the averages, and some explanation is

necessary to indicate their actual value.

In the feeding of leaves in 1914, there were three cases, two becoming sick. The third case received 0.661 pound without symptoms, the same quantity that was received by No. 282, which became sick. It seems probable, then, that the average figure 0.6615 must be pretty close to the toxic limit. In the feeding of "leaves and some young buds" in 1914, while the minimum of sick cases received 0.499 pound, another animal received 0.551 pound without ill effect; it is evident that the toxic limit must be not far from 0.5 pound. In the feeding of eight cases on "leaves, stems, flowers, and buds," with material collected near the station, the toxic limit was practically the same as in the preceding cases. A perusal of Table II shows that during the growth of the seed heads the toxicity was reduced and that the fully developed seed heads were somewhat less toxic than the plant in the earlier stages.

In the two cases of forced feeding of seeds in 1914 the average toxic dose was 0.153 pound; inasmuch as the animal receiving the maximum amount was very sick, the actual toxic limit must be considered to be close to the minimum figure of 0.11 pound. It will be noticed that the cases of feeding of seeds in 1913 had practically the same average dose as the cases of forced feeding in 1914; this is accounted for by the fact that these animals ate the seed in such a short time that the results were similar to those from forced feeding.

It is interesting to note in Table III that the lethal dose is only

slightly larger than the toxic dose.

In transferring these results to the probable dosage when sheep are range fed, the feeding habits of the sheep must be taken into consideration. In the corrals the sheep do not, as a rule, eat as readily as when on the range. When the sheep in a band are grazing together, both imitation and jealousy affect the quantity of any plant which a sheep eats in a given time; so it is reasonable to suppose that if feed is short and Zygadenus fairly abundant, sheep may eat much more in a short time than they would under corral conditions. Under such circumstances, the dosage might approximate that of the drenching experiments. Therefore, it appears probable that animals feeding on the range might, because of the more rapid eating, be poisoned with much less than when in corrals.

EXPERIMENTS WITH HORSES AND CATTLE.

Three experiments were made of feeding Zygadenus to a horse, as shown in Table I (section D), Z. venenosus being used in two tests and Z. elegans in one. The smallest quantity fed in these experiments was 12.1 pounds per 1,000 pounds of weight, and the largest was 15 pounds. In each instance the feeding was extended through several days, and the feeding of 15 pounds extended through 6 days. If the quantity necessary to poison a horse should be in the same proportion to its weight as that required to poison a sheep, it would be, according to our dosage, at least between 11 and 12 pounds, and probably much more. It may be presumed, therefore, that in these cases the amount fed was not sufficient to produce toxic effects. There is, however, abundance of evidence that horses are poisoned by Zygadenus, although not ordinarily with fatal results.

Section C of Table I shows the results of feeding Zygadenus venenosus to three cattle. Two of these animals, both of which received leaves and flowers, showed symptoms of poisoning, one on 58 pounds per 1,000 pounds of weight and the other on 46.5 pounds per 1,000 pounds of weight. In these cattle, therefore, the average toxic dose was 52.25 pounds, which was fed in an average of $6\frac{1}{2}$ days. This compares fairly well with the results reached with

sheep and would indicate that the toxic dose for cattle, computed in terms of the weight of the animal, does not differ materially from the toxic dose for sheep.

COMPARATIVE TOXICITY OF DIFFERENT SPECIES OF ZYGADENUS.

In the course of the experiments, four species of Zygadenus were used, Z. venenosus, Z. elegans, Z. paniculatus, and Z. coloradensis, by far the greater part of the work being done with Z. venenosus. The number of experiments with Z. elegans and Z. paniculatus was very small, and the material, especially in the case of Z. paniculatus, had been shipped a long distance, so that there was some question of the water content of the plant. Apparently, however, Z. elegans and Z. paniculatus do not differ materially in toxicity from Z. venenosus. Z. coloradensis, however, produced no toxic effects whatever with the exception of slight symptoms in one sheep, although the plant was fed in quantities several times as great as the toxic dose of Z. venenosus.

It is evident that in the feeding of cattle with Zygadenus coloradensis at Mount Carbon in 1909, the results of which are given in Table I, the quantities fed were too small to produce results, even if the plants were as poisonous as Z. venenosus. In the experiment of 1910, however, a large quantity was fed, and sufficiently large quantities in single days to produce symptoms of poisoning if the plant were as toxic as Z. venenosus.

In this connection it should be added that Dr. C. L. Alsberg made a laboratory examination of the Colorado plants and found in them a very small quantity of alkaloid. It would appear, then, that the form which is identified by some botanists as Z. coloradensis contains the same toxic substance as the other form, but that this substance is present in so small a quantity that it is unlikely that it ever produces toxic effects on domestic animals. While it is not in the province of this paper to discuss the systematic relations of plants, it may be suggested that this difference of toxicity between Z. elegans and Z. coloradensis may indicate a valid specific distinction between these two forms which are so closely related that by some botanists they are considered identical.

DOES TOXICITY VARY WITH LOCALITY?

The collections of Zygadenus venenosus with which experiments were made were obtained at the "Station" (by which is understood the region within a radius of 2 miles of the station), at an elevation of about 4,050 feet; at "Greycliff," $2\frac{1}{2}$ to 3 miles distant from the station, at an elevation of about 3,920 feet; and at "Cabin Corral" and "George Hughes's" (locations from 4 to 7 miles from the station), at an elevation of something over 5,000 feet. Material of this species

was also used from Avery, Cal., collected at an elevation of 3,500 feet. The material of Z. elegans was from two localities, from near Red Lodge, Mont., at an elevation between 5,500 and 6,000 feet, and from the Fishlake National Forest, Utah, at an elevation of something over 9,000 feet. The Z. paniculatus material was collected near Ephraim, Utah, at an elevation between 5,500 and 6,000 feet. All the Z. coloradensis material was collected within 4 or 5 miles of the Mount Carbon station, at an elevation of something over 10,000 feet.

As has been stated already, the lack of toxic properties in Zygadenus coloradensis is assumed to be characteristic of the species. The experiments with Z. elegans and Z. paniculatus were few in number, and too much importance must not be attached to the results. Apparently, however, not only did they have, practically, the same toxicity as the Z. venenosus collected near the station, but there was no evident difference between the Z. elegans of Montana and that collected in Utah. The Z. venenosus, collected in California gave the same results as that produced by material from the neighborhood of the Greycliff station.

An entirely unexplained variation in toxicity was exhibited by material collected at Cabin Corral, about 5 miles from the station and at a greater elevation of about a thousand feet. When Table II was being compiled, it was noticed that the cases receiving "forced feeding of leaves stems, flowers, and buds" fell into two distinct divisions, one with an average dosage of 0.5495 pound and the other with an average of 1.789 pounds. This difference was so marked that the two sets were separated in the summary. In searching for some possible explanation of this difference, it was found that all the cases with the larger dosage were treated with material collected at Cabin Corral. Note was then made of the other items in this table which were collected in this locality, and a glance at the table will show that in the other cases the Cabin-Corral material showed much less toxicity. It has been entirely impossible to explain this difference. The number of cases would seem to make it certain that this result was not due to an error of experimentation. There are no local conditions to account for it. The George Hughes place, at which collections were made giving the same results as those produced by the station material, is situated at about the same distance from the camp as Cabin Corral, at about the same elevation, and it has the same soil conditions. The question of the correlation of variation in toxicity with changes in altitude was raised, but the experiments do not indicate any such relation. The fact that the Cabin-Corral material was less toxic is nevertheless substantiated, and it would appear that while Zygadenus venenosus, Z. elegans, and Z. paniculatus have ordinarily the same degree of toxicity wherever grown, there is a possibility of marked variation.

EFFECT OF REPEATED FEEDING IN PRODUCING IMMUNITY OR INCREASED SUSCEPTIBILITY.

During the course of the experimental work at Greycliff a number of sheep were treated with Zygadenus several times during the same season. It was important to decide whether a sheep after having been poisoned once was more or less likely to be affected a second time. A careful analysis of the results showed that no effect either of immunity or of increased susceptibility was produced. The fact that an animal had suffered from poisoning once neither lessened the effect of another dose, nor, on the other hand, was the sheep any more likely to suffer from a second experience.

REMEDIES.

Because of the heavy losses of sheep from Zygadenus poisoning it was deemed important to investigate thoroughly the possibility of finding some remedial measures to reduce the number of deaths. To this end a large number of experiments were made, as can be seen by an examination of the table giving the summarized account of the work.

It has been shown by Hunt that the poisonous principle of Zygadenus is excreted in the urine, and this has been verified by the authors in the cases of some of the sheep used in the Greycliff experiments. Hunt concluded that the logical remedy is some diuretic which will insure excretion rapid enough to prevent serious effects from the poisoning, and his experiments seem to substantiate this position. He also advised the use of permanganate of potash administered per os to destroy the alkaloid in the stomach.

The experimental work on remedies in 1912 was based upon these conclusions of Hunt. Later, a number of remedies were used in the hope that some method might be found sufficiently simple to be used under range conditions. This work was carried on for three summers in order to get the average of a considerable number of cases, and a brief statement of the results of the more important experiments follows.

CAFFEIN AND DIURETIN.

The conclusions reached by Hunt led to a series of experiments with caffein. In 1912, caffein sodiobenzoate was administered to five animals subcutaneously; in two of these cases potassium permanganate was also used, and in one tannic acid. It was evident that by the use of this drug the excretion of urine was increased, but the observers could not see that any marked improvement followed in the condition of the animals. In the summer of 1913, diuretin and caffein citrate were given per os in four cases, of which one died and three recovered. In these cases, as in those of the preceding year,

there was no evidence of any good result. All these experiments had been with single doses. In 1914 two animals were treated, one with two doses of 10 grains each of caffein sodiobenzoate administered subcutaneously, and one with three doses of 5 grains each. One of these animals died and one recovered, but in neither case could it be seen that the remedy was advantageous. It seemed to be clear that while caffein might be considered a logical remedy it failed in practical application.

STRYCHNIN.

Although the work of Hunt quite clearly indicated that strychnin was not beneficial, it seemed best to try a few experiments to see whether, by its stimulating effect, it might not aid in relieving the depression of the animals. Six cases were treated by subcutaneous injections. In two of these cases eserin was also used, and in one case gin. There was an apparently beneficial effect in one case, but a study of all fails to show any good results which could be fairly considered as due to strychnin.

ESERIN, EPSOM SALTS, LINSEED OIL.

With the idea that relief might be brought about by an increase in intestinal elimination, eserin was administered subcutaneously, and Epsom salts and linseed oil per os; no reduction of toxic symptoms could be seen.

CHARCOAL.

Dr. Sollman suggested to the writers that charcoal, by adsorption, might be beneficial. Three experiments were made with this, in one case combined with linseed oil. No beneficial results followed.

WHISKY AND DIGITALIS.

In some cases of extreme depression whisky seemed to have an effect in bridging over a period when death might otherwise have followed. The same thing is true of digitalis, which in one or two cases may have saved the life of the patient. Neither drug, however, had any marked effect. It can only be said that if the symptoms of the animal are carefully watched, times will be found when whisky or digitalis may be administered advantageously. Inasmuch as the life of the individual sheep is of small importance, these remedies are of little practical use.

POTASSIUM PERMANGANATE.

Especial interest attaches to the experiments with potassium permanganate, since it is the remedy that has been most commonly recommended for plant poisoning. The dosage advised for a mature sheep has been 5 to 9 grains. This was used at first in the experimental work, and when no beneficial results appeared it was increased

to 30 grains, but still with no evidence of a reduction of the toxic effect. In two cases, 15 grains were introduced directly into the rumen with no better results.

Because of this lack of success it was deemed best to try mixing the permanganate with the Zygadenus before administration to see if the alkaloid would be destroyed in vitro. On May 19, 1913, two sheep of equal weight, Nos. 184 and 191, were each drenched with 0.586 pound of Zygadenus venenosus in water. In the dose given to No. 191 there were dissolved 7½ grains of potassium permanganate and 7½ grains of aluminum sulphate. Both animals were sick, and there was no recognizable difference in the degree of illness. The experiment tended to show that the administration of the potassium permanganate was without any definite effect upon the toxicity of the plant, but a similar experiment on July 11, 1914, on sheep No. 253 showed quite clearly that the dosage of the former experiment was insufficient. In this case 0.441 pound of seed heads of Z. venenosus was mixed in water with 15 grains of potassium permanganate and 15 grains of aluminum sulphate and the mixture allowed to stand for 20 minutes before being administered. The sheep displayed no symptoms of poisoning, although other cases of the same date receiving the same quantity of Zygadenus, with no remedy, showed distinct symptoms. It seemed clear that a sufficient quantity of permanganate will diminish the toxicity of the plant, when mixed with it before administration. When given after symptoms of poisoning are exhibited, however, the remedy is of no value. This, too, has been demonstrated by practical experience upon the range. Potassium permanganate has been used by many sheep owners in Montana. and it is the almost universal testimony that it is worthless.

TANNIC ACID.

Sheep No. 206, on June 4, 1913, was drenched with 0.43 pound of Zygadenus tops, including leaves and flowers. To this drench were added three grams of tannic acid. The animal showed no signs of illness. On May 29 a sheep was made sick on 0.389 pound, and on May 30 one was made sick on 0.385 pound and one died on 0.384 pound. It seems fair to presume, therefore, that the tannic acid had been of benefit to sheep No. 206.

Sheep No. 210, on June 6, 1913, was drenched with 0.496 pound of leaves and flowers of *Zygadenus venenosus* to which 3 grams of tannic acid had been added. On the same date sheep No. 209 received 0.495 pound of the same material, but without the tannic acid. Both animals were sick and recovered, but it was the impression of the observers that sheep No. 210 was not as sick as sheep No. 209.

On June 7, 1913, sheep Nos. 212 and 213 were each drenched with 0.55 pound of leaves and flowers of Zygadenus venenosus. In the

drench given to No. 212 were included 4 grams of tannic acid. This sheep had no symptoms of illness, while No. 213 died 1 hour and 17 minutes after the administration of the drench. These two sheep were of very nearly equal weight and the dose was the same (0.55 pound) per hundredweight of animal. There was every reason to expect similar results except for the effect of the tannic acid. Difference of individual susceptibility would seem to be eliminated in this instance, in which one animal died and the other showed no symptoms of poisoning.

During the season of 1913, four animals which had been fed on Zygadenus venenosus were given doses of tannic acid after toxic symptoms were well developed. All of these animals recovered. These cases, however, were not connected up with control cases, and it is possible that all would have recovered without any remedial aid.

The general result of all the experiments in 1913 with tannic acid indicated that it can be used with beneficial results. The experiments seemed also to indicate very clearly that, in vitro, the tannic acid was much more effective than potassium permaganate as an antidote for the Zygadenus alkaloid.

In 1914, a large number of cases were treated with tannic acid, in order to try it out thoroughly. In most of these experiments one or more control animals were used. Where the tannic acid was administered in a single dose, in 19 cases, there were only two deaths; in most of these cases, however, the Zygadenus was not given in a quantity necessarily fatal.

A study of the cases in which there was a control shows apparently beneficial results in some instances. For example, sheep Nos. 249 and 251 received the same quantity of Zygadenus on July 9; No. 249 died, while No. 251, which received a dose of tannic acid, lived.

Sheep Nos. 229 and 235 were fed the same quantity of Zygadenus on June 3. No. 229 was treated with tannic acid and was not so sick as No. 235. On the other hand, Nos. 239 and 256 were fed on June 16 with the same quantity, and No. 239, which received the tannic acid, had more marked symptoms than No. 256. Sheep Nos. 269, 255, and 282 were fed the same quantity of Zygadenus on June 15 and June 16. Tannic acid was administered to Nos. 269 and 282; both of these animals were sick, while No. 255 exhibited no symptoms. A consideration of all these cases shows that tannic acid in single doses can not be considered an effective remedy, although under favorable conditions some cases may be benefited.

SODIUM BICARBONATE.

It was suggested by Mr. O. F. Black that, inasmuch as alkaloids are, to a large extent, insoluble in an alkaline solution, sodium bicarbonate might serve to prevent the solution and absorption of the

poisonous principle of Zygadenus and thus prove valuable as a medicinal remedy. This was used only in repeated doses, and the results will be discussed under the next head.

REPEATED DOSES OF TANNIC ACID AND SODIUM BICARBONATE.

Inasmuch as tannic acid is a recognized remedial agent for poisoning by alkaloids, it seemed strange that so little benefit followed its use. In seeking for an explanation, it occurred to the writers that it might be accounted for by the fact that, because of the character of a ruminant's stomachs, the remedy does not actually come in contact with any considerable quantity of the poisonous substance. The first stomach of a ruminant always contains a large quantity of material. When an animal feeds upon a poisonous plant, the material taken up goes to the first stomach; some of this, after maceration, proceeds to the third and fourth stomachs, while another part goes on only after rumination. If the remedy is given in the form of a drench, it will be distributed in all the stomachs, although ordinarily the larger part of the drench goes directly to the third and fourth stomachs. That part of the drench which goes to the fourth stomach, we can assume, takes effect on the alkaloid which has arrived at that part of the digestive canal. The portion of the drench which stops in the first stomach meets a mass of organic matter, in which it is lost; there is no reason to think that any antidote for an alkaloid will have any selective effect, so as to attack the Zygadenus alkaloid rather than the multitude of other substances in the stomach with which it can unite. The only hope of destroying the alkaloid under such circumstances would be by flooding the first stomach with the antidote, and that is practically impossible. So even when the antidote is introduced by a canula directly into the first stomach, it would be impracticable to use a quantity sufficient to produce any marked effect.

On the other hand, inasmuch as no absorption takes place in the stomachs, if the antidote could meet the poisonous material as it passes through the fourth stomach good results might be expected. On the basis of this conclusion, it seemed best to the writers to try the effect of antidotes repeated at frequent intervals; it was thought that if the antidote could reach the fourth stomach frequently enough to catch the alkaloid as it passed from the first stomach and render it more or less innocuous before passing into the intestine, the remedy might be distinctly beneficial.

Four experiments of this character were conducted with tannic acid, all with controls, which received no remedy but were fed with the same quantity of Zygadenus. The tannic acid was given in doses of 1 and 2 grams, repeated at intervals varying from 10 to 30 minutes, or longer in some cases, at the latter part of the experiment.

The total time of treatment varied from 4 to $7\frac{1}{2}$ hours, and the total quantity of tannic acid given varied from 14 to 16 grams. The doses and intervals were as follows:

Sheep No. 263: 9 doses, 1 gram each, once in 10 minutes; 2 doses, 1 gram each, once in 30 minutes; 1 dose, 1 gram, in 20 minutes; 2 doses, 1 gram each, once in 30 minutes. Total, 14 grams.

Sheep No. 216: 7 doses, 2 grams each, once in 30 minutes. Total, 14 grams.

Sheep No. 267: 5 doses, 1 gram each, once in 10 minutes; 3 doses, 1 gram each, once in 15 minutes; 8 doses, 1 gram each, once in 30 minutes. Total, 16 grams.

Sheep No. 291: 3 doses, 2 grams each, once in 30 minutes; 1 dose, 2 grams, in 60 minutes; 1 dose, 2 grams, in 30 minutes; 2 doses, 2 grams each, once in 60 minutes; 1 dose, 2 grams, in 3 hours. Total, 16 grams.

All these animals recovered and were not as sick as the controls, Nos. 269 and 294. Sheep No. 291 suffered more than the others, but the tannic acid in this case was administered later in the illness, after a course of small doses of Epsom salts had failed to produce any effect. All these animals and the controls were given Zygadenus collected on the same date. The experiments were considered to prove conclusively that repeated doses of tannic acid are beneficial.

A similar set of experiments was conducted with sodium bicarbonate. Seven animals were used, and all, with one exception, were fed Zygadenus material collected on the same date, and in the exceptional case the material was collected only a few days later. Doses of sodium bicarbonate of 2 and 4 grams were given at intervals varying from 15 to 60 minutes. The total time of treatment was from $2\frac{1}{2}$ to 5 hours, and the total amount of sodium carbonate given varied from 20 to 48 grams. The doses and intervals were as follows:

Sheep No. 246: 10 doses, 4 grams each, once in 30 minutes. Total, 40 grams.

Sheep No. 259: 8 doses, 4 grams each, once in 15 minutes; 4 doses, 4 grams each, once in 30 minutes. Total, 48 grams.

Sheep No. 264: 6 doses, 4 grams each, once in 60 minutes. Total, 24 grams.

Sheep No. 292: 10 doses, 4 grams each, once in 30 minutes. Total, 40 grams.

Sheep No. 293: 4 doses, 4 grams each, once in 30 minutes; 2 doses, 2 grams each, once in 30 minutes. Total, 20 grams.

Sheep No. 277: 5 doses, 4 grams each, once in 30 minutes; 2 doses, 2 grams each, once in 30 minutes; 1 dose, 2 grams, after $1\frac{1}{4}$ hours. Total, 26 grams.

Sheep No. 240: 3 doses, 8 grams each, once in 60 minutes; 2 doses, 4 grams each, once in 60 minutes. Total, 32 grams.

Of these animals all recovered but one, No. 264. This sheep received a total of 24 grams, given at hour intervals. No. 240 also received the remedy at hour intervals and recovered very slowly, being unable to stand on the morning after the poisonous dose had been given. All the other cases, except No. 277, recovered rather quickly. No. 277 was as slow as No. 240, although the doses of sodium bicarbonate were given frequently, and the total amount was

greater than that given to No. 264 and to No. 293, which recovered. If we exclude No. 277, it would appear clear that sodium carbonate given in sufficiently frequent doses is distinctly beneficial. The Zygadenus in the case of No. 277 was given in three doses, and it is possible that there was some accumulative effect, which may explain in part the slow recovery.

The general conclusion from the experiments with sodium bicarbonate is that if the remedy is given at frequent intervals it will prove distinctly beneficial. The dose should be 4 grams, and this should

be repeated as often as every 30 minutes.

These experiments with repeated doses of tannic acid and sodium bicarbonate were interesting from a theoretical standpoint and indicate a line of treatment which can be used successfully with valuable animals. It is evident, however, that remedies used in this way can not be recommended for the ordinary band of sheep, for the expense of the treatment would be greater than the value of the animals.

No experiments of repeated doses were made with potassium permanganate; but it is probable that it could be used successfully, although the general trend of the experimental work is to indicate that the potassium permanganate is not, as a remedy, so efficient as tannic acid and sodium bicarbonate.

BLEEDING.

It is customary among sheep herders to bleed sheep poisoned by Zygadenus, the favorite place being the angular artery and vein of the eye. Although there seems to be no logical reason for this practice, it seemed wise to try it, and three sheep were treated in this way; two of the three died, and no beneficial result appeared in any of the cases.

METHODS OF PREVENTING LOSSES.

The most obvious thing to do is, of course, to keep the animals from eating the plant. With this end in view, it is important that all herders should be taught to recognize Zygadenus. When the plant is in flower this is not at all difficult, but it has been a matter of surprise to find to what extent, among the herders and sheep owners, the plant is not known, even at this stage. Before flowering, its grasslike leaves are not so easily recognized, but there is no reason why a fairly intelligent man should not be taught to know it even then. If one knows the plant in the preflowering and flowering stages, he will readily recognize it in the later dried-up condition, when, it will be remembered, it is fully as dangerous as earlier in the season.

When the plant is recognized care should be taken that the sheep do not have an opportunity to eat any large quantity of it. If it be necessary to drive the sheep over a patch of Zygadenus, the herder should take the precaution to have the band well fed before making the drive. If hungry sheep come upon a thick growth of Zygadenus, some of them, in their haste to satisfy their hunger, are almost certain to become poisoned, while if already well fed they are likely to choose their food with more care and to eat less of the Zygadenus.

Special care should be used early in the season, not because the plant is more poisonous at that time, but because, on account of the dry condition of other forage, it is more likely to be eaten. Later in the season sheep are less likely to eat a large quantity, because of the greater abundance of other food. As a matter of fact, most of the cases of extensive poisoning have occurred before the flowering of the plant.

If sheep become poisoned, they should be kept as quiet as possible. Any attempt to make them move about is likely to have disastrous

results.

So far as remedies are concerned, none has been found so far that gives much promise of being really useful. The experimental work at Greycliff shows that repeated doses of tannic acid or sodium bicarbonate will aid in recovery, but this method of treatment is not practically possible for animals upon the range.

GENERAL SUMMARY.

Zygadenus grows abundantly on many of the stock ranges of the West and is one of the most important sources of loss to sheepmen. Apparently all species of Zygadenus are poisonous. The plants are poisonous through the whole season of their growth, but the tops are somewhat more poisonous at the time of flowering. The toxicity of the bulbs and tops is about the same, while the seeds are much more toxic than other parts of the plant. Cases of poisoning are more likely to occur before the maturity of the plant, because at that time other forage is scanty.

The toxic dose varies according to the conditions of feeding. In drenched animals it may be put at about one-half a pound for an animal weighing a hundred pounds. In fed animals it varied from

1.6 pounds to 5.6 pounds.

The poisonous principle is an alkaloid or alkaloids allied to veratrin and cevadin.

Sheep, cattle, and horses are poisoned by the plant, but the fatalities are almost entirely confined to sheep.

The principal symptoms are salivation, nausea, muscular weakness, coma, and sometimes attacks of dyspnœa.

To prevent losses, it is important to recognize the plant and avoid grazing upon it. If animals become sick they should be kept quiet, and under this treatment many will recover. There is no satisfactory medical remedy.

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